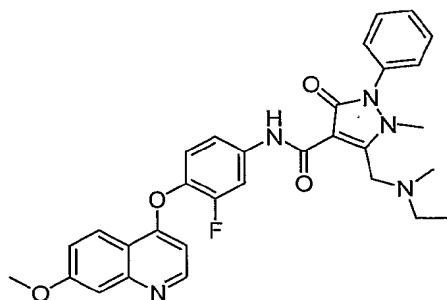
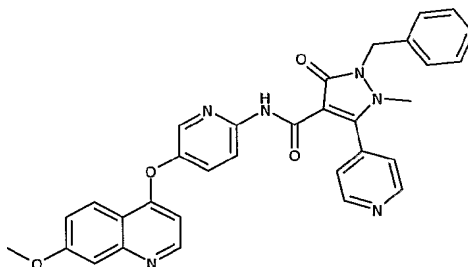


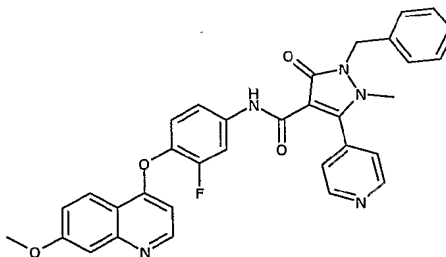
Hz, 1H), 3.75 – 3.64 (m, 3H), 2.51 (s, 3H), 2.01 – 1.91 (m, 1H), 1.84 (quintet, J = 6.0 Hz, 2H), 1.56 – 1.47 (m, 1H).

**Example 10**

- 5 **5-((ethyl(methyl)amino)methyl)-N-(3-fluoro-4-(7-methoxyquinolin-4-yloxy)phenyl)-1-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) m/z: 556 (MH<sup>+</sup>). Calc'd exact mass for C<sub>31</sub>H<sub>30</sub>FN<sub>5</sub>O<sub>4</sub>: 555. <sup>1</sup>H NMR (400 MHz, CDCl<sub>3</sub>): 11.06 (s, 1H), 8.59 (d, J = 5.0 Hz, 1H), 8.28 (d, J = 9.0 Hz, 1H), 7.92 (d, J = 12.0 Hz, 1H), 7.59 (t, J = 7.0 Hz, 2H), 7.53 – 7.48 (m, 1H), 7.43 – 7.36 (m, 3H), 7.31 (d, J = 9.0 Hz, 1H), 7.23 (d, J = 10.0 Hz, 1H), 7.17 (t, J = 9.0 Hz, 1H), 6.41 (d, J = 5.0 Hz, 1H), 4.22 (s, 2H), 3.98 (s, 3H), 3.58 (s, 3H), 2.63 (quartet, J = 7.0 Hz, 2H), 2.37 (s, 3H), 1.15 (t, J = 7.0 Hz, 3H).
- 10

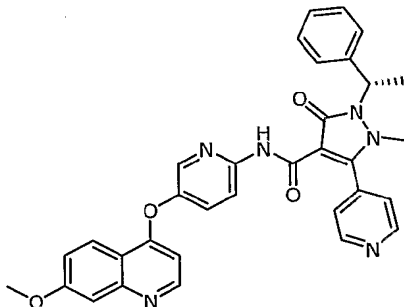
**Example 11**

- 2-benzyl-N-(5-(7-methoxyquinolin-4-yloxy)pyridin-2-yl)-1-methyl-3-oxo-5-(pyridin-4-yl)-2,3-dihydro-1H-pyrazole-4-carboxamide MS (ESI pos. ion) m/z 559 (MH<sup>+</sup>). Calc'd exact mass for C<sub>32</sub>H<sub>26</sub>N<sub>6</sub>O<sub>4</sub> 558.
- 15

**Example 12**

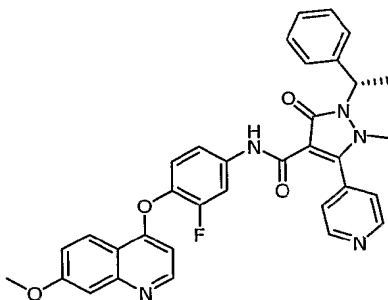
**2-benzyl-N-(3-fluoro-4-(7-methoxyquinolin-4-yloxy)phenyl)-1-methyl-3-oxo-5-(pyridin-4-yl)-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)  $m/z$  576 (MH<sup>+</sup>). Calc'd exact mass for C<sub>33</sub>H<sub>26</sub>FN<sub>5</sub>O<sub>4</sub> 575.

### Example 13



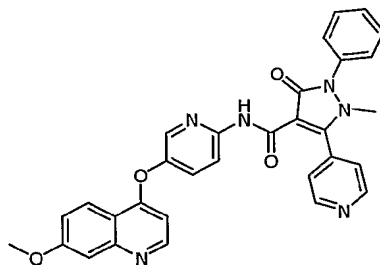
**(S)-N-(5-(7-methoxyquinolin-4-yloxy)pyridin-2-yl)-1-methyl-3-oxo-2-(1-phenylethyl)-5-(pyridin-4-yl)-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)  $m/z$ : 573 (MH<sup>+</sup>). Calc'd exact mass for C<sub>33</sub>H<sub>28</sub>N<sub>6</sub>O<sub>4</sub>: 572. <sup>1</sup>H NMR (400 MHz, CDCl<sub>3</sub>): 11.52 (s, 1H), 8.98 (d, J = 6.0 Hz, 2H), 8.84 (d, J = 6.0 Hz, 1H), 8.41 (d, J = 9.0 Hz, 1H), 8.37 (d, J = 3.0 Hz, 1H), 8.31 (d, J = 9.0 Hz, 1H), 7.81 (d, J = 5.0 Hz, 2H), 7.78 (d, J = 2.0 Hz, 1H), 7.60 (dd, J = 9.0 Hz, 2.0 Hz, 1H), 7.50 – 7.38 (m, 6H), 6.76 (d, J = 7.0 Hz, 1H), 6.17 (quartet, J = 8.0 Hz, 1H), 4.08 (s, 3H), 3.13 (s, 3H), 2.02 (d, J = 7.0 Hz, 3H).

### Example 14

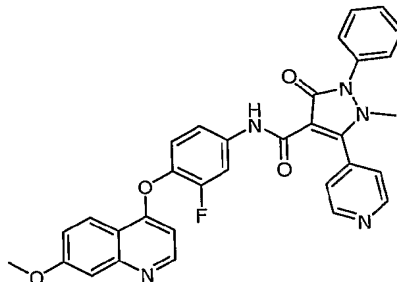


**(S)-N-(3-fluoro-4-(7-methoxyquinolin-4-yloxy)phenyl)-1-methyl-3-oxo-2-(1-phenylethyl)-5-(pyridin-4-yl)-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)  $m/z$ : 590 (MH<sup>+</sup>). Calc'd exact mass for C<sub>34</sub>H<sub>28</sub>FN<sub>5</sub>O<sub>4</sub>: 589.

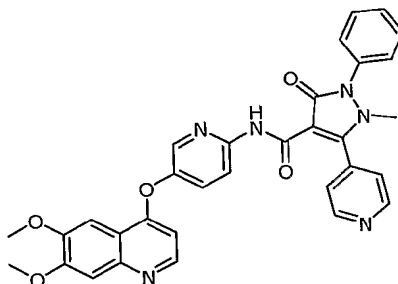
### Example 15



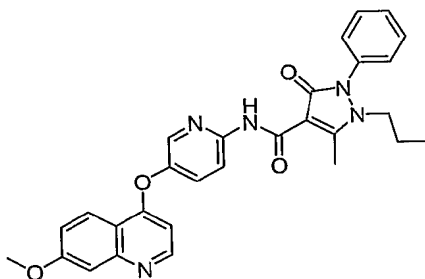
**N-(5-(7-methoxyquinolin-4-yloxy)pyridin-2-yl)-1-methyl-3-oxo-2-phenyl-5-(pyridin-4-yl)-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) m/z: 545 (MH<sup>+</sup>). Calc'd exact mass for C<sub>31</sub>H<sub>24</sub>N<sub>6</sub>O<sub>4</sub>: 544.

**Example 16**

**N-(3-fluoro-4-(7-methoxyquinolin-4-yloxy)phenyl)-1-methyl-3-oxo-2-phenyl-5-(pyridin-4-yl)-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) m/z: 562 (MH<sup>+</sup>). Calc'd exact mass for C<sub>32</sub>H<sub>24</sub>FN<sub>5</sub>O<sub>4</sub>: 561.

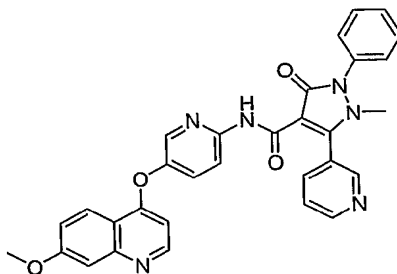
**Example 17**

**N-(5-(6,7-dimethoxyquinolin-4-yloxy)pyridin-2-yl)-1-methyl-3-oxo-2-phenyl-5-(pyridin-4-yl)-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) m/z: 575 (MH<sup>+</sup>). Calc'd exact mass for C<sub>32</sub>H<sub>26</sub>N<sub>6</sub>O<sub>5</sub>: 574.

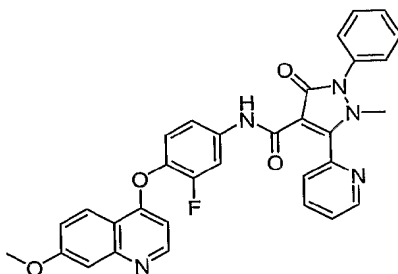
**Example 18**

**N-(5-(7-methoxyquinolin-4-yloxy)pyridin-2-yl)-5-methyl-3-oxo-2-phenyl-1-propyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) m/z: 510 (MH<sup>+</sup>). Calc'd exact mass for C<sub>29</sub>H<sub>27</sub>N<sub>5</sub>O<sub>4</sub>: 509. <sup>1</sup>HNMR (300 MHz, CDCl<sub>3</sub>): 0.81 (t, 3H), 1.54 (m, 2H), 2.82 (s,

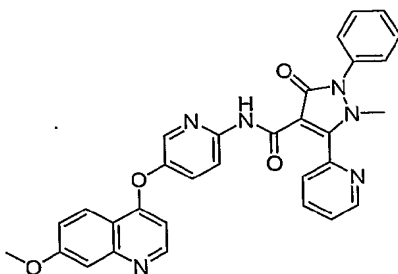
3H), 3.75 (m, 2H), 4.0 (s, 3H), 6.45 (d, 1H), 7.31 (d, 2H), 7.41-7.52 (m, 6H), 8.30 (d, 2H), 8.40 (d, 1H), 8.60 (d, 1H).

**Example 19**

5 **N-(5-(7-methoxyquinolin-4-yloxy)pyridin-2-yl)-1-methyl-3-oxo-2-phenyl-5-(pyridin-3-yl)-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) m/z: 545 (MH<sup>+</sup>). Calc'd exact mass for C<sub>31</sub>H<sub>24</sub>N<sub>6</sub>O<sub>4</sub>: 544. <sup>1</sup>HNMR (300 MHz, CDCl<sub>3</sub>): 3.42 (s, 3H), 3.89 (s, 3H), 4.37-4.49 (m, 1H), 5.22 (s, 1H), 6.34 (d, J=5.26 Hz, 1H), 7.15 (dd, J=9.21, 2.48 Hz, 1H), 7.20 (s, 2H), 7.28-7.50 (m, 7H), 8.15 (d, J=6.14 Hz, 1H), 8.17 (s, 1H), 8.29 (d, J=9.06 Hz, 1H), 8.52 (d, J=5.26, 1H), 11.55 (s, 1H).

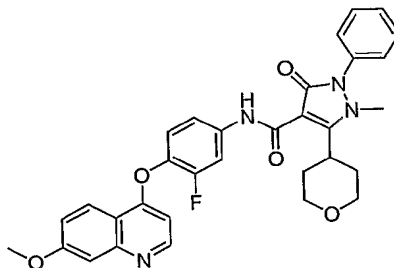
**Example 20**

15 **N-(3-fluoro-4-(7-methoxyquinolin-4-yloxy)phenyl)-1-methyl-3-oxo-2-phenyl-5-(pyridin-2-yl)-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) m/z: 562 (MH<sup>+</sup>). Calc'd exact mass for C<sub>32</sub>H<sub>24</sub>FN<sub>5</sub>O<sub>4</sub>: 561. <sup>1</sup>HNMR (300 MHz, CDCl<sub>3</sub>): 2.80 (s, 1H), 3.34 (s, 3H), 3.96 (s, 3H), 6.39 (d, J=5.12 Hz, 1H), 7.11-7.32 (m, 4H), 7.40-7.64 (m, 6H), 7.79-8.01 (m, 3H), 8.25 (d, J=9.21 Hz, 1H), 8.57 (d, J=5.26 Hz, 1H), 8.80 (d, J=4.53 Hz, 1H), 11.12 (s, 1H).

**Example 21**

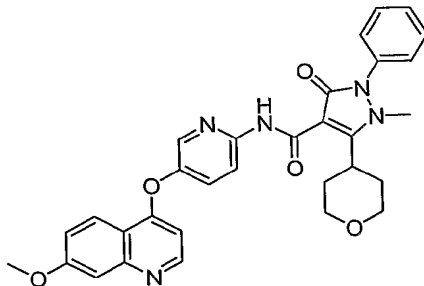
**N-(5-(7-methoxyquinolin-4-yloxy)pyridin-2-yl)-1-methyl-3-oxo-2-phenyl-5-(pyridin-2-yl)-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) m/z: 545 (MH<sup>+</sup>). Calc'd exact mass for C<sub>31</sub>H<sub>24</sub>N<sub>6</sub>O<sub>4</sub>: 544. <sup>1</sup>HNMR (300 MHz, CDCl<sub>3</sub>): 3.24 (s, 3H), 3.88 (s, 3H), 6.32 (d, J=5.26 Hz, 1H), 7.13 (dd, J=9.13, 2.41 Hz, 1H), 7.20 (s, 1H), 7.31-7.53 (m, 7H), 7.80-7.90 (m, 2H), 8.10-8.23 (m, 3H), 8.50 (d, J=5.26 Hz, 1H), 8.72 (d, J=4.82 Hz, 1H), 11.39 (s, 1H).

### Example 22



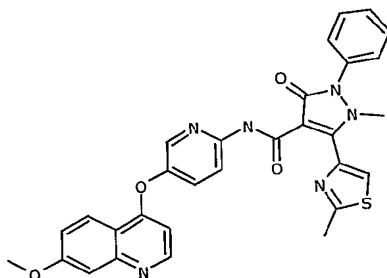
**N-(3-fluoro-4-(7-methoxyquinolin-4-yloxy)phenyl)-1-methyl-3-oxo-2-phenyl-5-(tetrahydro-2H-pyran-4-yl)-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) m/z: 569 (MH<sup>+</sup>). Calc'd exact mass for C<sub>32</sub>H<sub>29</sub>FN<sub>4</sub>O<sub>5</sub>: 568. <sup>1</sup>HNMR (300 MHz, CDCl<sub>3</sub>): 1.83 (d, J=10.96 Hz, 2H), 2.33 (qd, J=12.57, 4.38 Hz, 2H), 3.49 (s, 3H), 3.63 (t, J=11.03 Hz, 2H), 3.96 (s, 3H), 4.15 (dd, J=11.40, 3.80 Hz, 2H), 4.47-4.60 (m, 1H), 5.29 (s, 1H), 6.40 (dd, J=5.19, 0.80 Hz, 1H), 7.14-7.42 (m, 4H), 7.46-7.60 (m, 3H), 7.92 (dd, J=12.50, 2.27 Hz, 1H), 8.27 (d, J=9.21 Hz, 1H), 8.59 (d, J=5.26 Hz, 1H), 11.25 (s, 1H).

### Example 23



**N-(5-(7-methoxyquinolin-4-yloxy)pyridin-2-yl)-1-methyl-3-oxo-2-phenyl-5-(tetrahydro-2H-pyran-4-yl)-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) m/z: 552 (MH<sup>+</sup>). Calc'd exact mass for C<sub>31</sub>H<sub>29</sub>N<sub>5</sub>O<sub>5</sub>: 551. <sup>1</sup>HNMR (300 MHz, CDCl<sub>3</sub>): 1.76 (d, J=10.67 Hz, 2H), 2.19-2.35 (m, 2H), 3.42 (s, 3H), 3.50-3.60 (m, 2H), 3.89 (s, 3H), 4.07 (dd, J=11.55, 3.80 Hz, 2H), 4.37-4.49 (m, 1H), 5.22 (s, 1H), 6.34 (d, J=5.26 Hz, 1H), 7.15 (dd, J=9.21, 2.48 Hz, 1H), 7.28-7.50 (m, 5H), 8.15 (d, J=6.14 Hz, 1H), 8.17 (s, 1H), 8.29 (d, J=9.06 Hz, 1H), 8.52 (d, J=5.26 Hz, 1H), 11.55 (s, 1H).

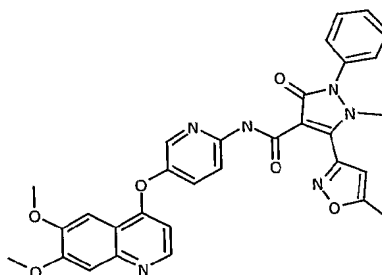
## Example 24



**1-Methyl-N-(5-((7-(methoxy)-4-quinolinyl)oxy)-2-pyridinyl)-5-(2-methyl-1,3-thiazol-4-yl)-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) m/z: 565

(MH<sup>+</sup>). Calc'd exact mass for C<sub>30</sub>H<sub>24</sub>N<sub>6</sub>O<sub>4</sub>S: 564. <sup>1</sup>H NMR (400 MHz, DMSO-*d*<sub>6</sub>) 11.51 (s, 1 H), 8.62 (d, *J*=5.18 Hz, 1 H), 8.32 - 8.36 (m, 2 H), 8.29 (d, *J*=9.09 Hz, 1 H), 8.22 (d, *J*=9.09 Hz, 1 H), 7.77 (dd, *J*=9.16, 2.84 Hz, 1 H), 7.61 - 7.68 (m, 2 H), 7.53 - 7.60 (m, 3 H), 7.42 (d, *J*=2.40 Hz, 1 H), 7.30 (dd, *J*=9.09, 2.40 Hz, 1 H), 6.53 (d, *J*=5.18 Hz, 1 H), 3.94 (s, 3 H), 3.28 - 3.33 (m, 3 H), 2.78 (s, 3 H):

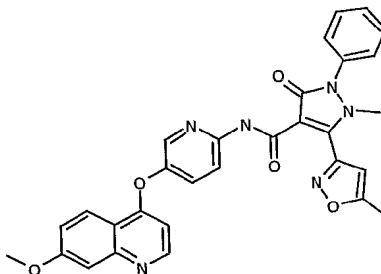
## Example 25



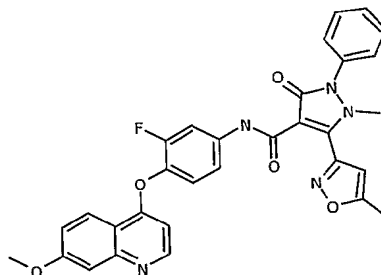
**N-(5-((6,7-bis(methoxy)-4-quinolinyl)oxy)-2-pyridinyl)-1-methyl-5-(5-methyl-3-isoxazolyl)-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)

m/z: 579 (MH<sup>+</sup>). Calc'd exact mass for C<sub>31</sub>H<sub>26</sub>N<sub>6</sub>O<sub>6</sub>: 578.

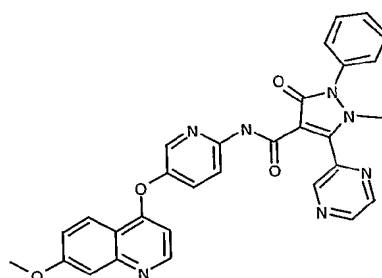
## Example 26



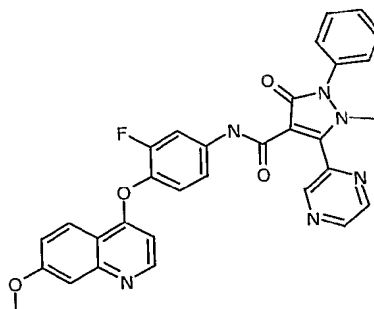
**1-methyl-5-(5-methyl-3-isoxazolyl)-N-(5-((7-(methyloxy)-4-quinolinyloxy)-2-pyridinyl)-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) m/z: 549 (MH<sup>+</sup>). Calc'd exact mass for C<sub>30</sub>H<sub>24</sub>N<sub>6</sub>O<sub>5</sub>: 548.

**Example 27**

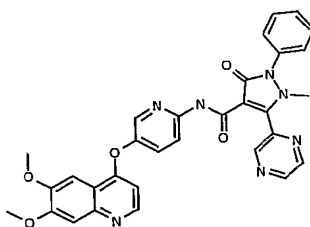
**N-(3-fluoro-4-((7-(methyloxy)-4-quinolinyloxy)phenyl)-1-methyl-5-(5-methyl-3-isoxazolyl)-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) m/z: 566 (MH<sup>+</sup>). Calc'd exact mass for C<sub>31</sub>H<sub>24</sub>FN<sub>5</sub>O<sub>5</sub>: 565.

**Example 28**

**1-methyl-N-(5-((7-(methyloxy)-4-quinolinyloxy)-2-pyridinyl)-3-oxo-2-phenyl-5-(2-pyrazinyl)-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) m/z: 546 (MH<sup>+</sup>). Calc'd exact mass for C<sub>30</sub>H<sub>23</sub>N<sub>7</sub>O<sub>4</sub>: 545.

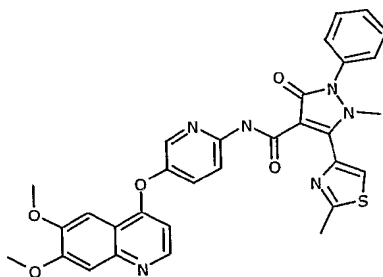
**Example 29**

**N-(3-fluoro-4-((7-(methyloxy)-4-quinolinyloxy)phenyl)-1-methyl-3-oxo-2-phenyl-5-(2-pyrazinyl)-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) m/z: 563.2 (MH<sup>+</sup>). Calc'd exact mass for C<sub>31</sub>H<sub>23</sub>FN<sub>6</sub>O<sub>4</sub>: 562.

**Example 30**

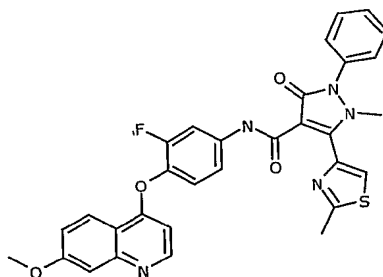
**N-(5-((6,7-bis(methoxy)-4-quinolinyloxy)-2-pyridinyl)-1-methyl-3-oxo-2-phenyl-5-(2-pyrazinyl)-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)  $m/z$ : 576 ( $MH^+$ ).

- 5 Calc'd exact mass for  $C_{31}H_{25}N_7O_5$ : 575.  $^1H$  NMR (400 MHz,  $DMSO-d_6$ ) 11.28 (s, 1 H), 9.11 (d,  $J=1.14$  Hz, 1 H), 8.87 - 8.90 (m, 1 H), 8.85 (d,  $J=2.53$  Hz, 1 H), 8.48 (d,  $J=5.31$  Hz, 1 H), 8.34 (d,  $J=2.91$  Hz, 1 H), 8.22 (d,  $J=9.09$  Hz, 1 H), 7.74 (dd,  $J=8.97, 2.91$  Hz, 1 H), 7.64 - 7.69 (m, 2 H), 7.57 - 7.63 (m, 3 H), 7.52 (s, 1 H), 7.40 (s, 1 H), 6.52 (d,  $J=5.31$  Hz, 1 H), 3.94 (d,  $J=5.31$  Hz, 6 H), 3.30 (s, 3H).

**Example 31**

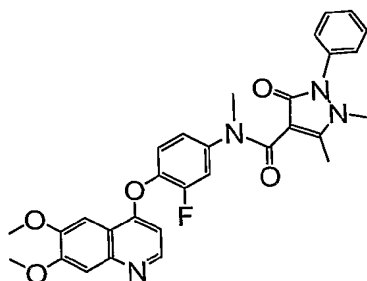
**N-(5-((6,7-bis(methoxy)-4-quinolinyloxy)-2-pyridinyl)-1-methyl-5-(2-methyl-1,3-thiazol-4-yl)-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)

- 15  $m/z$ : 595 ( $MH^+$ ). Calc'd exact mass for  $C_{31}H_{26}N_6O_5S$ : 594.

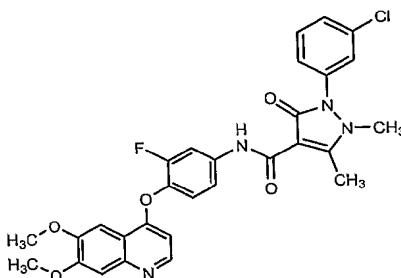
**Example 32**

**N-(3-fluoro-4-((7-(methoxy)-4-quinolinyloxy)phenyl)-1-methyl-5-(2-methyl-1,3-thiazol-4-yl)-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)  $m/z$ : 582

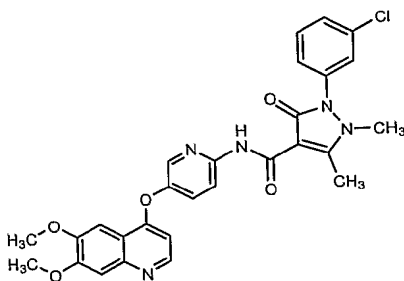
- 20 ( $MH^+$ ). Calc'd exact mass for  $C_{31}H_{24}FN_5O_4S$ : 581.

**Example 33**

**N-(4-(6,7-dimethoxyquinolin-4-yloxy)-3-fluorophenyl)-N,1,5-trimethyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)  $m/z$  543 (MH<sup>+</sup>) Calc'd exact mass for C<sub>30</sub>H<sub>27</sub>FN<sub>4</sub>O<sub>5</sub> 543. <sup>1</sup>H NMR (300 MHz, CDCl<sub>3</sub>) 8.28 (1 H, d, 5.3 Hz), 7.56 (1 H, s), 7.33 - 7.53 (4 H, m), 7.24 - 7.10 (5 H, m), 6.26 (1 H, d, =5.4 Hz), 4.06 (6 H, s), 3.52 (3 H, s), 3.22 (3 H, s), 2.54 (3 H, s).

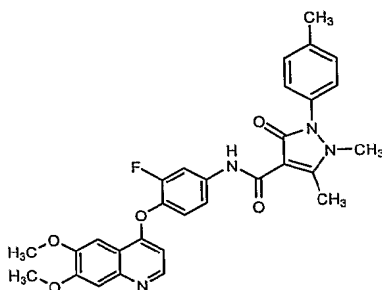
**Example 34**

**2-(3-chlorophenyl)-N-(4-(6,7-dimethoxyquinolin-4-yloxy)-3-fluorophenyl)-1,5-dimethyl-3-oxo-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)  $m/z$ : 563 (MH<sup>+</sup>). Calc'd exact mass for C<sub>29</sub>H<sub>24</sub>ClFN<sub>4</sub>O<sub>5</sub>: 562.

**Example 35**

**2-(3-chlorophenyl)-N-(5-(6,7-dimethoxyquinolin-4-yloxy)pyridin-2-yl)-1,5-dimethyl-3-oxo-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)  $m/z$ : 546 (MH<sup>+</sup>). Calc'd exact mass for C<sub>28</sub>H<sub>24</sub>ClN<sub>5</sub>O<sub>5</sub>: 545.

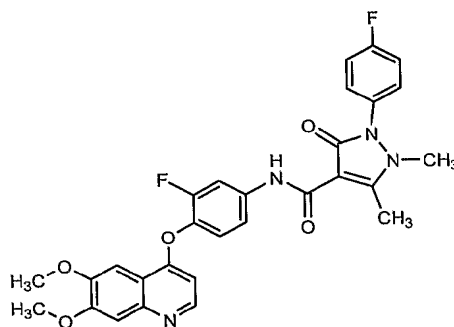
## Example 36



**N-(4-(6,7-dimethoxyquinolin-4-yloxy)-3-fluorophenyl)-1,5-dimethyl-3-oxo-2-p-tolyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) m/z: 543 (MH<sup>+</sup>). Calc'd exact

5 mass for C<sub>30</sub>H<sub>27</sub>FN<sub>4</sub>O<sub>5</sub>: 542. <sup>1</sup>H NMR (400 MHz, DMSO-*d*<sub>6</sub>) 11.00 (s, 1 H), 8.48 (d, *J*=5.31 Hz, 1 H), 7.98 (dd, *J*=13.20, 2.21 Hz, 1 H), 7.53 (s, 1 H), 7.29 - 7.45 (m, 7 H), 6.47 (d, *J*=5.43 Hz, 1 H), 3.95 (s, 6 H), 3.35 (s, 3 H), 2.70 (s, 3 H), 2.40 (s, 3 H).

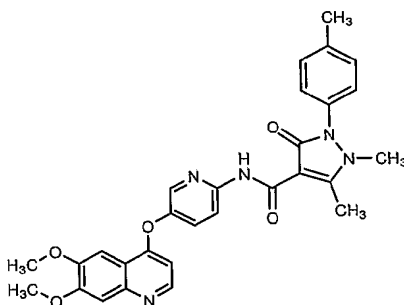
## Example 37



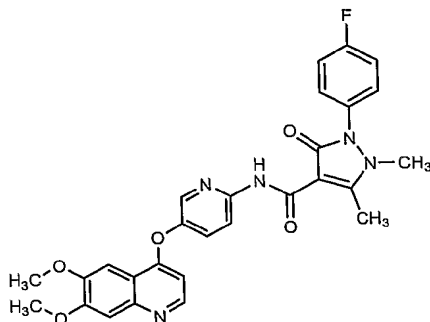
10 **N-(4-(6,7-dimethoxyquinolin-4-yloxy)-3-fluorophenyl)-2-(4-fluorophenyl)-1,5-dimethyl-3-oxo-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) m/z: 547 (MH<sup>+</sup>). Calc'd exact mass for C<sub>29</sub>H<sub>24</sub>F<sub>2</sub>N<sub>4</sub>O<sub>5</sub>: 546. <sup>1</sup>H NMR (400 MHz, DMSO-*d*<sub>6</sub>) 10.94 (s, 1 H), 8.48 (d, *J*=5.18 Hz, 1 H), 7.98 (dd, *J*=13.07, 2.34 Hz, 1 H), 7.32 - 7.55 (m, 8 H), 6.47 (d, *J*=5.05 Hz, 1 H), 3.95 (s, 6 H), 3.37 (s, 3 H), 2.70 (s, 3 H).

15

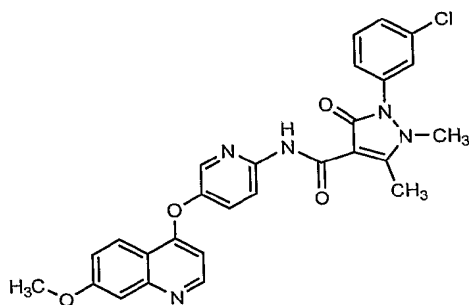
## Example 38



**N-(5-(6,7-dimethoxyquinolin-4-yloxy)pyridine-2-yl)-1,5-dimethyl-3-oxo-2-p-tolyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)  $m/z$ : 526 (MH<sup>+</sup>). Calc'd exact mass for C<sub>29</sub>H<sub>27</sub>N<sub>5</sub>O<sub>5</sub>: 525.

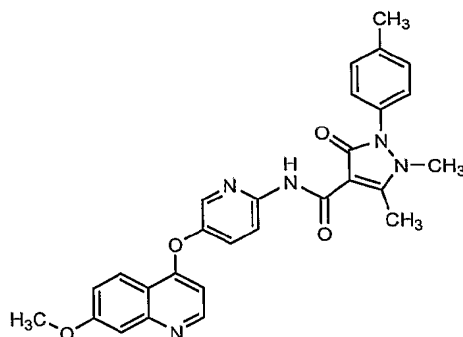
**Example 39**

**N-(5-(6,7-dimethoxyquinolin-4-yloxy)pyridin-2-yl)-2-(4-fluorophenyl)-1,5-dimethyl-3-oxo-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)  $m/z$ : 530 (MH<sup>+</sup>). Calc'd exact mass for C<sub>28</sub>H<sub>24</sub>FN<sub>5</sub>O<sub>5</sub>: 529.

**Example 40**

**2-(3-chlorophenyl)-N-(5-(7-methoxyquinolin-4-yloxy)pyridin-2-yl)-1,5-dimethyl-3-oxo-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)  $m/z$ : 516 (MH<sup>+</sup>). Calc'd exact mass for C<sub>27</sub>H<sub>22</sub>ClN<sub>5</sub>O<sub>4</sub>: 515. <sup>1</sup>H NMR (400 MHz, DMSO-*d*<sub>6</sub>) 11.15 (s, 1 H), 8.62 (d, *J*=5.30 Hz, 1 H), 8.32 - 8.39 (m, 2 H), 8.22 (d, *J*=9.22 Hz, 1 H), 7.81 (dd, *J*=8.97, 3.03 Hz, 1 H), 7.57 - 7.66 (m, 3 H), 7.41 - 7.46 (m, 2 H), 7.30 (dd, *J*=9.16, 2.59 Hz, 1 H), 6.54 (d, *J*=5.18 Hz, 1 H), 3.94 (s, 3 H), 3.40 (s, 3 H), 2.73 (s, 3 H).

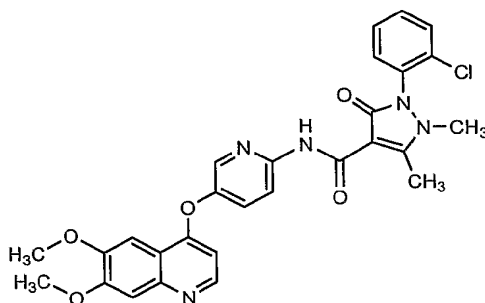
## Example 41



**N-(5-(7-methoxyquinolin-4-yloxy)pyridin-2-yl)-1,5-dimethyl-3-oxo-2-p-tolyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)  $m/z$ : 496 (MH<sup>+</sup>). Calc'd exact mass for

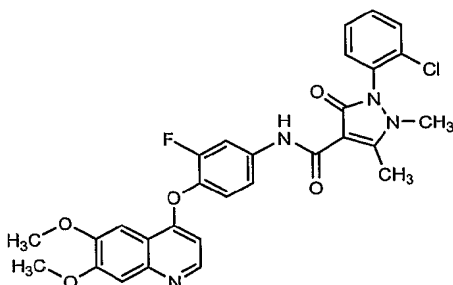
- 5  $C_{28}H_{25}N_5O_4$ : 495. <sup>1</sup>H NMR (400 MHz, DMSO-*d*<sub>6</sub>) 11.29 (s, 1 H), 8.62 (d,  $J=4.93$  Hz, 1 H), 8.30 - 8.40 (m, 2 H), 8.22 (d,  $J=9.22$  Hz, 1 H), 7.80 (d,  $J=9.60$  Hz, 1 H), 7.26 - 7.46 (m, 6 H), 6.53 (d,  $J=4.93$  Hz, 1 H), 3.94 (s, 3 H), 3.35 (s, 3 H), 2.71 (s, 3 H), 2.40 (s, 3 H).

## Example 42

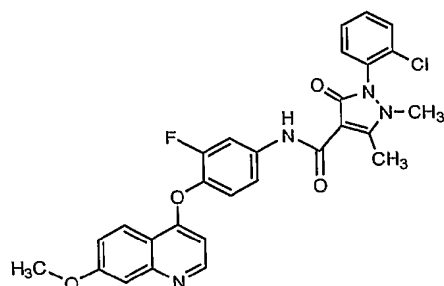


- 10 **2-(2-chlorophenyl)-N-(5-(6,7-dimethoxyquinolin-4-yloxy)pyridin-2-yl)-1,5-dimethyl-3-oxo-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)  $m/z$ : 546 (MH<sup>+</sup>). Calc'd exact mass for  $C_{28}H_{24}ClN_5O_5$ : 545.

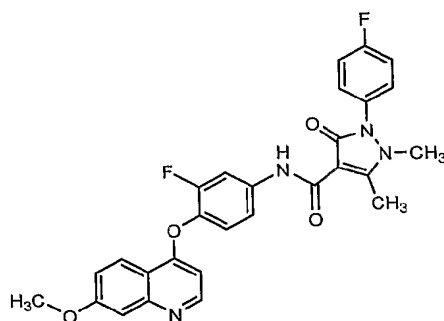
## Example 43



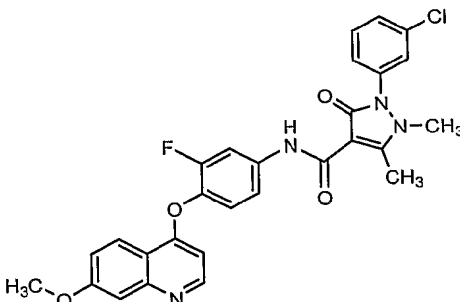
- 15 **2-(2-chlorophenyl)-N-(4-(6,7-dimethoxyquinolin-4-yloxy)-3-fluorophenyl)-1,5-dimethyl-3-oxo-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)  $m/z$ : 563 (MH<sup>+</sup>). Calc'd exact mass for  $C_{29}H_{24}ClFN_4O_5$ : 562.

**Example 44**

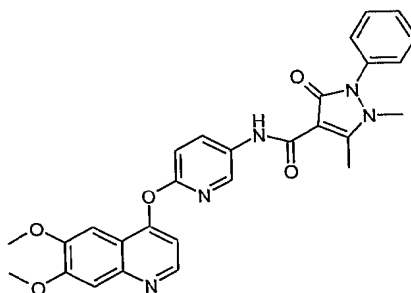
**2-(2-chlorophenyl)-N-(3-fluoro-4-(7-methoxyquinolin-4-yloxy)phenyl)-1,5-dimethyl-3-oxo-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)  $m/z$ : 533 (MH<sup>+</sup>). Calc'd exact mass for C<sub>28</sub>H<sub>22</sub>ClFN<sub>4</sub>O<sub>4</sub>: 532. <sup>1</sup>H NMR (400 MHz, DMSO-*d*<sub>6</sub>) 10.88 (s, 1 H), 8.61 (d,  $J=5.18$  Hz, 1 H), 8.23 (d,  $J=9.09$  Hz, 1 H), 7.99 (dd,  $J=13.07$ , 1.71 Hz, 1 H), 7.77 (d,  $J=7.71$  Hz, 1 H), 7.55 - 7.72 (m, 3 H), 7.37 - 7.47 (m, 2 H), 7.26 - 7.38 (m, 2 H), 6.48 (d,  $J=5.18$  Hz, 1 H), 3.94 (s, 3 H), 3.34 (s, 3 H), 2.71 (s, 3 H).

**Example 45**

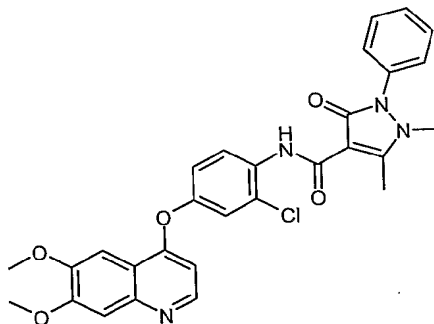
**N-(3-fluoro-4-(7-methoxyquinolin-4-yloxy)phenyl)-2-(4-fluorophenyl)-1,5-dimethyl-3-oxo-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)  $m/z$ : 517 (MH<sup>+</sup>). Calc'd exact mass for C<sub>28</sub>H<sub>22</sub>F<sub>2</sub>N<sub>4</sub>O<sub>4</sub>: 516. <sup>1</sup>H NMR (400 MHz, DMSO-*d*<sub>6</sub>) 10.95 (s, 1 H), 8.61 (d,  $J=5.18$  Hz, 1 H), 8.23 (d,  $J=9.09$  Hz, 1 H), 7.99 (dd,  $J=12.88$ , 1.89 Hz, 1 H), 7.27 - 7.57 (m, 8 H), 6.48 (d,  $J=5.18$  Hz, 1 H), 3.94 (s, 3 H), 3.36 (s, 3 H), 2.70 (s, 3 H).

**Example 46**

**2-(3-chlorophenyl)-N-(3-fluoro-4-(7-methoxyquinolin-4-yloxy)phenyl)-1,5-dimethyl-3-oxo-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)  $m/z$ : 533 (MH<sup>+</sup>). Calc'd exact mass for C<sub>28</sub>H<sub>22</sub>ClFN<sub>4</sub>O<sub>4</sub>: 532. <sup>1</sup>H NMR (400 MHz, DMSO-*d*<sub>6</sub>) 10.85 (s, 1 H), 8.62 (d,  $J=5.18$  Hz, 1 H), 8.24 (d,  $J=9.09$  Hz, 1 H), 7.99 (dd,  $J=13.01, 2.40$  Hz, 1 H), 7.57 - 7.66 (m, 3 H), 7.29 - 7.47 (m, 5 H), 6.49 (d,  $J=5.05$  Hz, 1 H), 3.95 (s, 3 H), 3.41 (s, 3 H), 2.72 (s, 3 H).

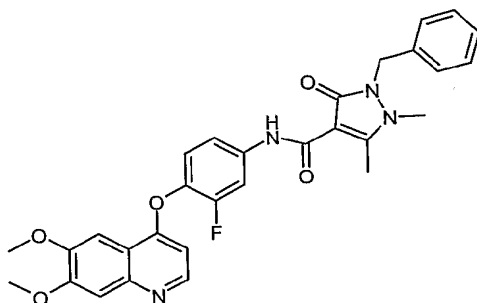
**Example 47**

**N-(6-(6,7-dimethoxyquinolin-4-yloxy)pyridin-3-yl)-1,5-dimethyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)  $m/z$ : 512 (MH<sup>+</sup>). Calc'd exact mass for C<sub>28</sub>H<sub>25</sub>N<sub>5</sub>O<sub>5</sub>: 511 <sup>1</sup>H NMR (400 MHz, DMSO-*d*<sub>6</sub>) 10.86 (s, 1 H), 8.56 (d,  $J=5.05$  Hz, 1 H), 8.50 (d,  $J=2.65$  Hz, 1 H), 8.27 (dd,  $J=2.9, 5.81$  Hz, 1H), 7.7-7.38 (3m, 7 H), 7.31 (d,  $J=8.72$  Hz, 1 H), 6.84 (d,  $J=5.18$  Hz, 1H), 3.95 (s, 3 H), 3.89 (s, 3 H), 3.38 (s, 3H), 2.71 (s, 3 H).

**Example 48**

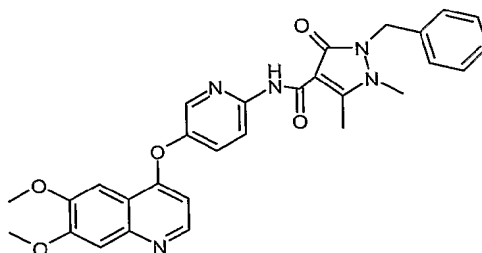
**N-(2-chloro-4-(6,7-dimethoxyquinolin-4-yloxy)phenyl)-1,5-dimethyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)  $m/z$ : 545 (MH<sup>+</sup>). Calc'd exact mass for C<sub>29</sub>H<sub>25</sub>ClN<sub>4</sub>O<sub>5</sub>: 544 <sup>1</sup>H NMR (400 MHz, CDCl<sub>3</sub>) 11.12 (s, 1 H), 8.70 (d,  $J=8.79$  Hz, 1 H), 8.50 (d,  $J=5.37$  Hz, 1H), 7.43 - 7.63 (m, 5 H), 7.38 (d,  $J=7.32$  Hz, 2H), 7.11 (dd,  $J=9.28, 1.95$  Hz, 1 H), 6.55 (d,  $J=3.91$  Hz, 1 H), 4.07 (s, 3 H), 4.04 (s, 3 H), 3.48 (s, 1 H), 3.37 (s, 3 H), 2.81 (s, 3 H).

## Example 49



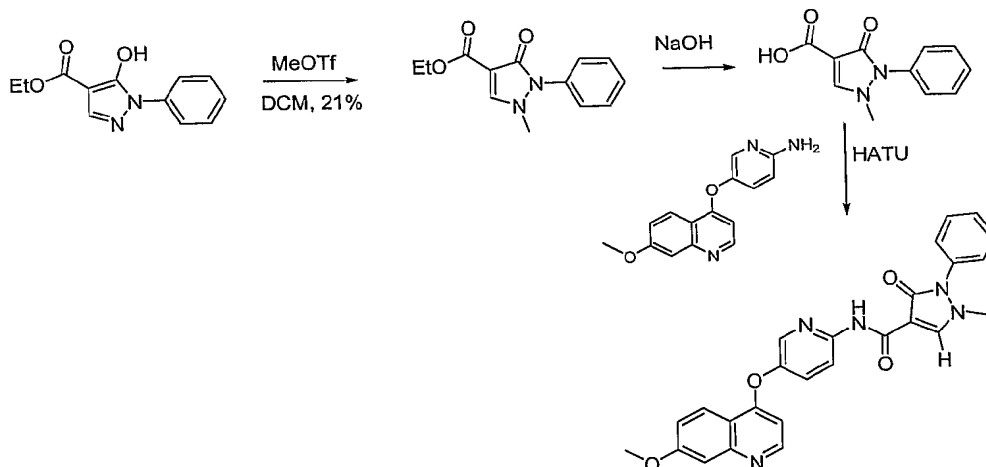
**2-benzyl-N-(4-(6,7-dimethoxyquinolin-4-yloxy)-3-fluorophenyl)-1,5-dimethyl-3-oxo-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)  $m/z$ : 543 (MH<sup>+</sup>). Calc'd exact mass for C<sub>30</sub>H<sub>27</sub>FN<sub>4</sub>O<sub>5</sub>: 542. 1H NMR (400 MHz, CDCl<sub>3</sub>) 10.98 (s, 1 H), 8.50 (d,  $J=5.37$  Hz, 1 H), 7.97 (d,  $J=12.70$  Hz, 1H), 7.61 (s, 1 H), 7.53 (s, 1 H), 7.42 - 7.29 (m, 4 H), 7.24 - 7.15 (m, 3 H), 6.49 (s, 1 H), 5.18 (s, 2 H), 4.07 (s, 6 H), 3.40 (s, 3 H), 2.67 (s, 3H)

## Example 50

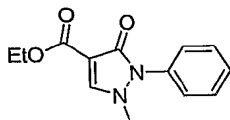


**2-benzyl-N-(5-(6,7-dimethoxyquinolin-4-yloxy)pyridin-2-yl)-1,5-dimethyl-3-oxo-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)  $m/z$ : 526 (MH<sup>+</sup>). Calc'd exact mass for C<sub>29</sub>H<sub>27</sub>N<sub>5</sub>O<sub>5</sub>: 525. 1H NMR (400 MHz, CDCl<sub>3</sub>) 11.43 (s, 1 H), 8.51 (d,  $J=5.37$  Hz, 1H), 8.40 (d,  $J=8.79$  Hz, 1 H), 8.29 (s, 1 H), 7.67 - 7.12 (m, 8 H), 6.53 - 6.44 (m, 1 H), 5.18 (s, 2 H), 4.07 (s, 3 H), 3.40 (s, 6 H), 2.66 (s, 3 H)

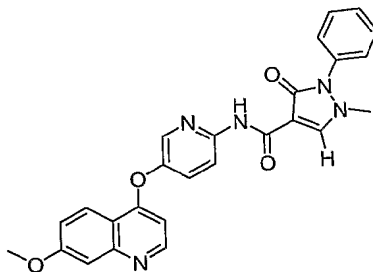
## Example 51



**N-(5-(7-methoxyquinolin-4-yloxy)pyridin-2-yl)-1-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide.**



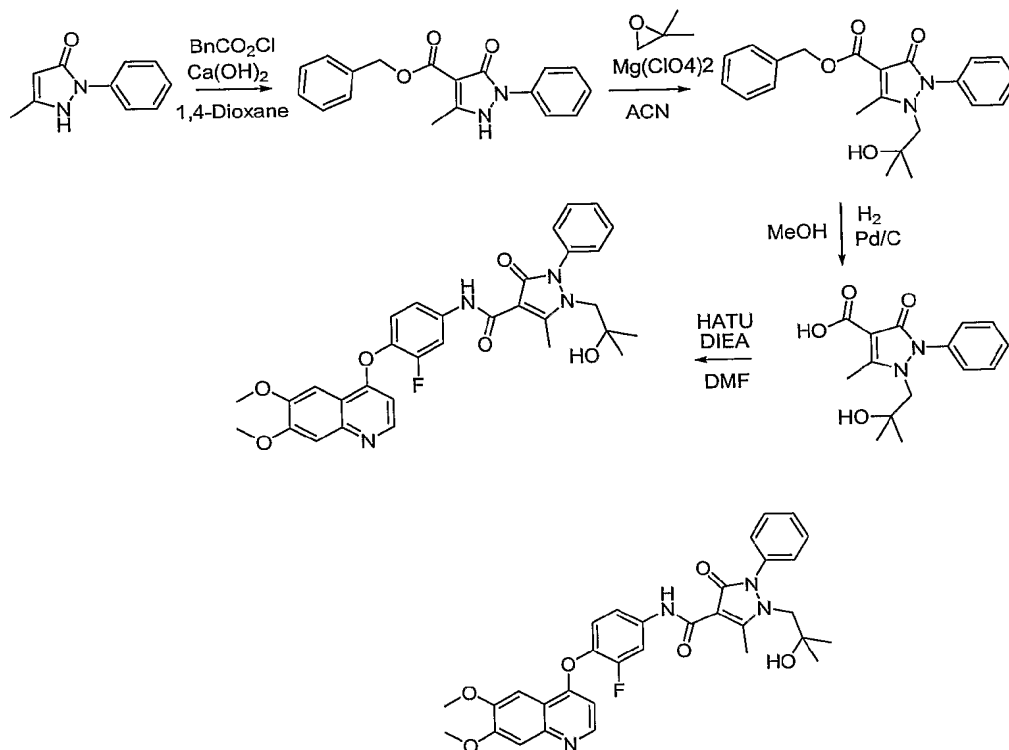
**Step 1: Ethyl 1-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxylate.** To a solution of ethyl 3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxylate (1000 mg, 5.0 mmol) in dichloromethane (10 mL) was added methyl trifluoromethanesulfonate (1200 mg, 7.3 mmol). The red solution was stirred at room temperature. After 14 h, the mixture was partitioned between dichloromethane and NaHCO<sub>3</sub> (sat). The aqueous was extracted with dichloromethane (2x). The combined organic was dried over Na<sub>2</sub>SO<sub>4</sub>, concentrated and purified on silica. The product was triturated with EtOAc-hexane-CHCl<sub>3</sub> to give the pure product as crystals (260 mg, 21%). Calc'd for C<sub>12</sub>H<sub>12</sub>N<sub>2</sub>O<sub>3</sub>, 232.08; MS (ESI pos. ion) m/z: 233 (MH<sup>+</sup>). <sup>1</sup>H NMR (400 MHz, CHLOROFORM-d): 1.36 (t, J=7.04 Hz, 3 H), 3.39 (s, 3 H), 4.32 (q, J=7.17 Hz, 2 H), 7.32 (d, J=7.43 Hz, 2 H), 7.42 (t, J=7.34 Hz, 1 H), 7.50 (t, J=7.73 Hz, 2 H), 7.99 (s, 1 H).



**Step 2: N-(5-(7-methoxyquinolin-4-yloxy)pyridin-2-yl)-1-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide.** A solution of ethyl 1-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxylate (260 mg, 1056 μmol) in MeOH was treated with NaOH (1000 μl, 5000 μmol) in H<sub>2</sub>O (3 mL). The mixture was heated to 60°C for 30 min and then cooled to room temperature. Then, the mixture was neutralized with aq. HCl (5 N, 1.1 mL) and concentrated to dryness. The residue was further dried with (azeotrope distillation with toluene, 3 x 5 mL). The resulting carboxylic acid was mixed with 5-(7-methoxyquinolin-4-yloxy)pyridin-2-amine (282 mg, 1054 μmol), Et<sub>3</sub>N (500 μl, 3587 μmol), and HATU (401 mg, 1054 μmol) in DMF (4 mL) - dichloromethane (5 mL) and was stirred at 60°C for 2 h. Upon cooling to room temperature, the mixture was diluted with EtOAc containing 10% MeOH (30 mL) and washed with H<sub>2</sub>O. The organic layer was dried over Na<sub>2</sub>SO<sub>4</sub>, concentrated, and eluted on silica (1-10% 2N NH<sub>3</sub>-MeOH in CHCl<sub>3</sub>). The product was further purified on preparative HPLC to afford a white powder (100 mg, 20%).

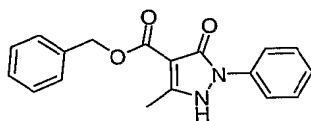
Calc'd for  $C_{26}H_{21}N_5O_4$ : 467.16; MS (ESI pos. ion)  $m/z$ : 468 (MH<sup>+</sup>). <sup>1</sup>H NMR (400 MHz, DMSO- $d_6$ ) 3.49 (s, 3 H) 3.95 (s, 3 H) 6.55 (d, J 5.1, 1 H) 7.30 (dd, J 2.0, 9.0, 1 H) 7.42 (s, 1 H) 7.59 (s, 17 H) 7.50- 7.60 (m, 5 H), 7.84 (dd, J 2.8, 9.2, 1H), 8.22 (d, J 9.2, 1H), 8.34 - 8.38 (m, 2 H) 8.62 (d, J 5.3, 1 H) 8.69 (s, 1 H) 10.86 (s, 1 H).

5

**Example 52**

**N-(4-(6,7-dimethoxyquinolin-4-yloxy)-3-fluorophenyl)-1-(2-hydroxy-2-methylpropyl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide**

10

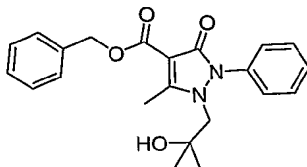


**Step 1: Benzyl 5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxylate.** A

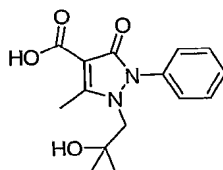
suspension of 3-methyl-1-phenyl-1H-pyrazol-5-ol (10.0 g, 57 mmol) and calcium hydroxide (8.5 g, 115 mmol) in dry 1,4-dioxane (100mL) was heated to 50 °C for 20 min. The

15 suspension was chilled to 10 °C and benzyl chloroformate (8.2 ml, 57 mmol) was in dioxane (10mL) added. The resultant was heated to 90 °C for 3h, cooled to 25 °C and then, chilled (0°C) 1 M HCl (200 mL) was added. The mixture was stirred at 25 °C overnight. A solid collected by filtration was washed with cold EtOH (2x25mL) and ether (50mL), dried at 80°C (sand bath) exposed to air for 4h to give the title compound (14.0g, 79% yield) as an off-white

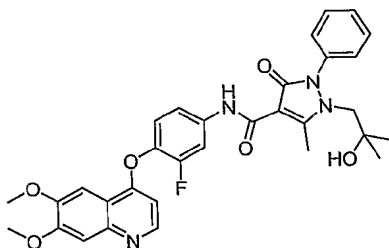
20 solid. MS (ESI pos. ion)  $m/z$ : 309 (MH<sup>+</sup>). Calc'd exact mass for  $C_{18}H_{16}N_2O_3$  308.



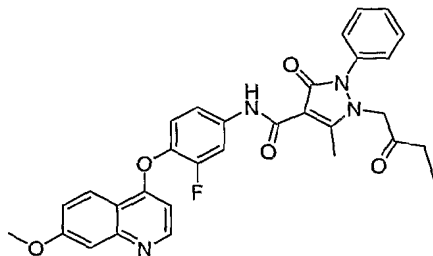
**Step 2: Benzyl 1-(2-hydroxy-2-methylpropyl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxylate.** To a stirring suspension of benzyl 5-hydroxy-3-methyl-1-phenyl-1H-pyrazole-4-carboxylate (3380 mg, 11 mmol) in chlorobenzene (30mL) at 10 °C under nitrogen was added trimethylaluminum (16 mL, 33 mmol, 2M in toluene). Internal temperature reached 27 °C. At 25°C, 1,2-epoxy-2-methylpropane (1000 mg, 16 mmol) was added. The reaction mixture was stirred for 3 h at 25°C, and then diluted with THF (500mL). The resultant was chilled to 10°C, and sodium sulfate decahydrate (2g) was added. After 1hr, another 2g of sodium sulfate decahydrate added. After 2h, the gel was filtered through a bed of celite and washed with EtOAc (3x100mL). The filtrate was then washed with aq. 1M HCl (50mL) and brine. The organic layer was dried over MgSO<sub>4</sub> and concentrated. The residue was purified on 120g silica chromatography (30>90% EtOAc/hex). To give the title compound (1.11g, 27% yield) as an amorphous solid. MS (ESI pos. ion) m/z: 381 (MH<sup>+</sup>). Calc'd exact mass for C<sub>22</sub>H<sub>24</sub>N<sub>2</sub>O<sub>4</sub> 380.



**Step 3: 1-(2-hydroxy-2-methylpropyl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxylic acid.** To a stirring solution of benzyl 1-(2-hydroxy-2-methylpropyl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxylate (300 mg, 789 μmol) in MeOH (10mL) was purged with argon for 10min. To this solution was added Pd/C (40mg), and the mixture was stirred for 3h under balloon of hydrogen. Reaction was monitored by LCMS. The reaction mixture was filtered through a bed of celite and concentrated under reduced pressure to give the title compound (220 mg, 96.1% yield) as an off-white solid. MS (ESI pos. ion) m/z: 291 (MH<sup>+</sup>). Calc'd exact mass for C<sub>15</sub>H<sub>18</sub>N<sub>2</sub>O<sub>4</sub> 290.

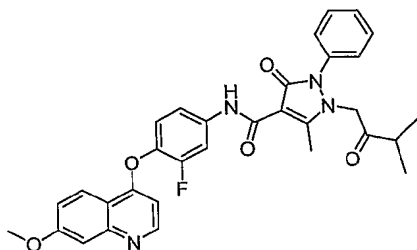


**Step 4: N-(4-(6,7-dimethoxyquinolin-4-yloxy)-3-fluorophenyl)-1-(2-hydroxy-2-methylpropyl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide.** To a stirring solution of 1-(2-hydroxy-2-methylpropyl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxylic acid (104 mg, 358  $\mu$ mol) and diisopropylethylamine (62  $\mu$ l, 358  $\mu$ mol) in DMF (1mL) was added HATU (136 mg, 358  $\mu$ mol) and stirred at 37 °C under nitrogen for 15min. To this was added 3-fluoro-4-(7-methoxyquinolin-4-yloxy)benzenamine (98 mg, 344  $\mu$ mol) and stirred overnight at 37 °C. The reaction mixture was diluted with dichloromethane (10mL) and washed with 1M NaOH (20mL), and extracted with dichloromethane (3x5mL). Combined organics layer was washed with brine and then dried with MgSO<sub>4</sub>. Residual DMF was removed with repeated azeotroping with toluene (4x5mL) under reduced pressure. The residue was purified on 12g silica (10>30% of 6% 2M NH<sub>3</sub> in MeOH/DCM). Final material was lyophilized from 50% ACN/water to give the title compound (155mg, 74% yield) as a white fluffy solid. MS (ESI pos. ion) m/z: 587 (MH<sup>+</sup>). Calc'd exact mass for C<sub>32</sub>H<sub>31</sub>FN<sub>4</sub>O<sub>6</sub> 587. <sup>1</sup>H NMR (400 MHz, CHLOROFORM-d) 10.88 (1 H, s), 8.47 (1 H, d, J=5.3 Hz), 7.92 (1 H, dd, J=12.5, 2.0 Hz), 7.59 (1 H, s), 7.54 (2 H, t, J=7.7 Hz), 7.40 - 7.49 (2 H, m), 7.30 (3 H, d, J=7.8 Hz), 7.17 (1 H, t, J=8.7 Hz), 6.45 (1 H, d, J=5.3 Hz), 4.06 (3 H, s), 4.05 (3 H, s), 3.88 (2 H, s), 2.89 (3 H, s), 2.01 (1 H, s), 1.15 (6 H, s)

**Example 53**

**N-(3-fluoro-4-(7-methoxyquinolin-4-yloxy)phenyl)-5-methyl-3-oxo-1-(2-oxobutyl)-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) m/z: 555 (MH<sup>+</sup>). Calc'd exact mass for C<sub>31</sub>H<sub>27</sub>FN<sub>4</sub>O<sub>5</sub>: 554. <sup>1</sup>H NMR (400 MHz, CHLOROFORM-d) 10.86 (1 H, s), 8.60 (1 H, d, J=5.3 Hz), 8.28 (1 H, d, J=9.2 Hz), 7.65 - 8.00 (2 H, m), 7.41 - 7.58 (4 H, m), 7.28 - 7.35 (3 H, m), 7.23 (1 H, dd, J=9.2, 2.3 Hz), 7.17 (1 H, t, J=8.7 Hz), 6.44 (1 H, d, J=5.3 Hz), 4.51 (2 H, s), 3.97 (3 H, s), 2.68 (3 H, s), 2.26 (2 H, q, J=7.2 Hz), 1.01 (3 H, t, J=7.2 Hz).

**Example 54**

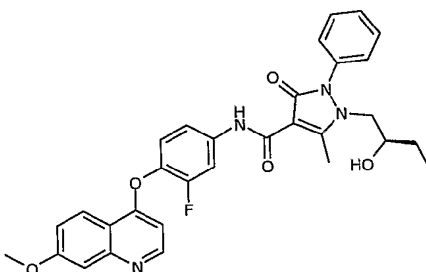


**N-(3-fluoro-4-(7-methoxyquinolin-4-yloxy)phenyl)-5-methyl-1-(3-methyl-2-oxobutyl)-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) m/z: 569

(MH<sup>+</sup>). Calc'd exact mass for C<sub>32</sub>H<sub>29</sub>FN<sub>4</sub>O<sub>5</sub>: 568. <sup>1</sup>H NMR (400 MHz, CHLOROFORM-d)

10.88 (1 H, s), 8.59 (1 H, d, J=5.3 Hz), 8.21 - 8.30 (2 H, m), 7.97 - 8.03 (1 H, m), 7.87 - 7.95 (2 H, m), 7.71 (1 H, t, J=7.4 Hz), 7.44 - 7.57 (3 H, m), 7.41 (1 H, d, J=2.2 Hz), 7.26 - 7.34 (3 H, m), 7.23 (1 H, dd, J=9.2, 2.3 Hz), 7.17 (1 H, t, J=8.7 Hz), 6.42 (1 H, d, J=5.3 Hz), 4.58 (2 H, s), 3.97 (3 H, s), 2.67 (3 H, s), 2.38 - 2.52 (1 H, m), 2.26 (3 H, s), 0.96 (6 H, d, J=6.8 Hz).

#### Example 55

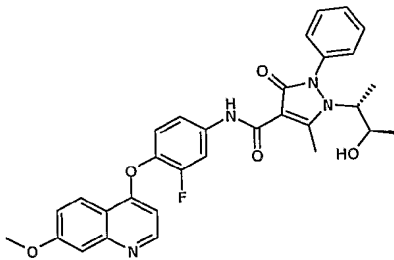


**(R)-N-(3-fluoro-4-(7-methoxyquinolin-4-yloxy)phenyl)-1-(2-hydroxybutyl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) m/z: 557

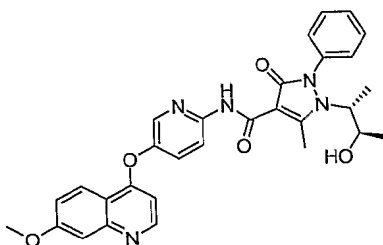
(MH<sup>+</sup>). Calc'd exact mass for C<sub>31</sub>H<sub>29</sub>FN<sub>4</sub>O<sub>5</sub>: 556. <sup>1</sup>H NMR (400 MHz, CHLOROFORM-d)

10.90 (1 H, s), 8.50 (1 H, d, J=5.3 Hz), 8.30 (1 H, d, J=9.2 Hz), 7.95 (1 H, dd, J=12.5, 2.0 Hz), 7.39 - 7.53 (3 H, m), 7.36 (1 H, d, J=2.2 Hz), 7.11 - 7.31 (6 H, m), 6.42 (3 H, d, J=5.3 Hz), 3.95 (3 H, s), 3.79 - 3.90 (1 H, m), 3.70 (1 H, dd, J=1.6 Hz), 3.56 - 3.66 (1 H, m), 2.85 (3 H, s), 1.60 - 1.82 (1 H, m), 1.27 - 1.43 (2 H, m), 0.83 (3 H, t, J=7.3 Hz)

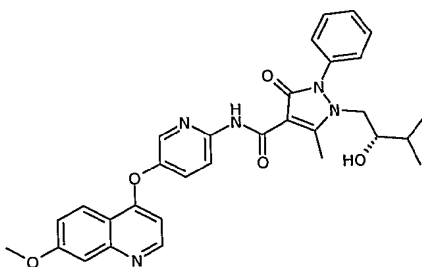
#### Example 56



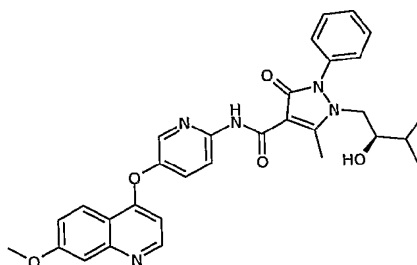
**N-(3-fluoro-4-(7-methoxyquinolin-4-yloxy)phenyl)-1-((2R,3R)-3-hydroxybutan-2-yl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) m/z: 557 (MH<sup>+</sup>). Calc'd exact mass for C<sub>31</sub>H<sub>29</sub>FN<sub>4</sub>O<sub>5</sub>: 556. <sup>1</sup>H NMR (400 MHz, CHLOROFORM-d) 10.94 (1 H, s), 8.54 (1 H, d, J=5.1 Hz), 8.27 (1 H, d, J=9.0 Hz), 7.90 (1 H, dd, J=12.3, 2.0 Hz), 7.43 - 7.57 (3 H, m), 7.37 - 7.40 (1 H, m), 7.31 (2 H, d, J=7.6 Hz), 7.25 - 7.28 (1 H, m), 7.23 (1 H, dd, J=9.2, 2.2 Hz), 7.16 (1 H, t, J=8.7 Hz), 6.40 (1 H, d, J=5.1 Hz), 3.77 - 4.05 (6 H, m), 2.89 (3 H, s), 1.63 - 1.91 (2 H, m), 1.50 (3 H, d, J=7.0 Hz), 1.11 (3 H, d, J=6.3 Hz)

**Example 57**

**1-((2R,3R)-3-hydroxybutan-2-yl)-N-(5-(7-methoxyquinolin-4-yloxy)pyridin-2-yl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) m/z: 540 (MH<sup>+</sup>). Calc'd exact mass for C<sub>30</sub>H<sub>29</sub>N<sub>5</sub>O<sub>5</sub>: 539. <sup>1</sup>H NMR (400 MHz, CHLOROFORM-d) 11.30 (1 H, s), 8.59 (1 H, d, J=5.3 Hz), 8.36 (1 H, d, J=9.0 Hz), 8.21 - 8.26 (2 H, m), 7.43 - 7.56 (4 H, m), 7.41 (1 H, d, J=2.5 Hz), 7.30 - 7.35 (2 H, m), 7.23 (1 H, dd, J=9.2, 2.5 Hz), 6.42 (1 H, d, J=5.3 Hz), 3.97 - 4.05 (1 H, m), 3.97 (3 H, s), 3.84 - 3.93 (1 H, m), 3.49 (1 H, s), 2.90 (3 H, s), 1.50 (3 H, d, J=7.2 Hz), 1.12 (3 H, d, J=6.3 Hz)

**Example 58**

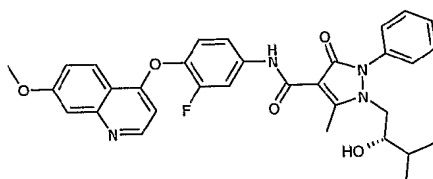
**(S)-1-(2-hydroxy-3-methylbutyl)-N-(5-(7-methoxyquinolin-4-yloxy)pyridin-2-yl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) m/z: 554 (MH<sup>+</sup>). Calc'd exact mass for C<sub>31</sub>H<sub>31</sub>N<sub>5</sub>O<sub>5</sub>: 553. <sup>1</sup>H NMR (400 MHz, DMSO-d<sub>6</sub>) 11.29 (1 H, s), 8.63 (1 H, d, J=5.1 Hz), 8.37 (1 H, d, J=9.0 Hz), 8.32 (1 H, d, J=2.9 Hz), 8.22 (1 H, d, J=9.2 Hz), 7.80 (1 H, dd, J=9.0, 2.9 Hz), 7.60 (2 H, t, J=7.5 Hz), 7.51 (1 H, t, J=7.3 Hz), 7.40 - 7.47 (3 H, m), 7.30 (1 H, dd, J=9.2, 2.5 Hz), 6.54 (1 H, d, J=5.3 Hz), 5.10 (1 H, d, J=5.9 Hz), 3.87 - 3.98 (4 H, m), 3.71 (1 H, d), 3.10 - 3.21 (1 H, m), 2.76 (3 H, s), 1.33 - 1.46 (1 H, m), 0.64 (3 H, d, J=6.8 Hz), 0.58 (3 H, d, J=6.8 Hz)

**Example 59**

**(R)-1-(2-hydroxy-3-methylbutyl)-N-(5-(7-methoxyquinolin-4-yloxy)pyridin-2-yl)-5-**

**methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) m/z:

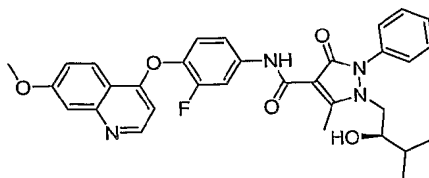
5 554 (MH<sup>+</sup>). Calc'd exact mass for C<sub>31</sub>H<sub>31</sub>N<sub>5</sub>O<sub>5</sub>: 553. <sup>1</sup>H NMR (400 MHz, DMSO-d<sub>6</sub>) 11.30 (1 H, s), 8.63 (1 H, d, J=5.1 Hz), 8.37 (1 H, d, J=9.0 Hz), 8.33 (1 H, d, J=2.7 Hz), 8.22 (1 H, d, J=9.2 Hz), 7.80 (1 H, dd, J=9.1, 2.8 Hz), 7.60 (2 H, t, J=7.5 Hz), 7.51 (1 H, t, J=7.3 Hz), 7.39 - 7.47 (3 H, m), 7.30 (1 H, dd, J=9.2, 2.5 Hz), 6.54 (1 H, d, J=5.1 Hz), 5.11 (1 H, d, J=5.9 Hz), 3.85 - 3.97 (4 H, m), 3.71 (2 H, s), 3.10 - 3.21 (2 H, m), 2.76 (3 H, s), 1.32 - 1.47 (1 H, m),  
10 0.64 (3 H, d, J=6.7 Hz), 0.58 (3 H, d, J=6.8 Hz)

**Example 60**

**(S)-N-(3-fluoro-4-(7-methoxyquinolin-4-yloxy)phenyl)-1-(2-hydroxy-3-methylbutyl)-5-**

**methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) m/z:

15 571 (MH<sup>+</sup>). Calc'd exact mass for C<sub>32</sub>H<sub>31</sub>FN<sub>4</sub>O<sub>5</sub>: 570. <sup>1</sup>H NMR (400 MHz, CHLOROFORM-d) 10.93 (1 H, s), 8.47 (1 H, d, J=5.3 Hz), 8.31 (1 H, d, J=9.2 Hz), 7.89 - 8.03 (1 H, m), 7.39 - 7.50 (3 H, m), 7.34 (1 H, d, J=2.0 Hz), 7.19 - 7.31 (3 H, m), 7.08 (2 H, d, J=7.2 Hz), 6.45 (1 H, d, J=5.3 Hz), 3.95 (3 H, s), 3.86 (1 H, dd, J=14.9 Hz), 3.64 - 3.77 (1 H, m), 3.44 - 3.48 (4 H, m), 3.33 - 3.42 (1 H, m), 2.82 (9 H, s), 1.46 - 1.61 (3 H, m), 0.80 (3 H, d, J=6.7 Hz), 0.72 (3 H,  
20 d, J=6.8 Hz)

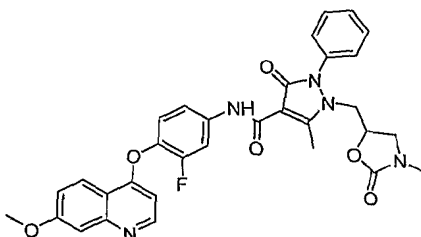
**Example 61**

**(R)-N-(3-fluoro-4-(7-methoxyquinolin-4-yloxy)phenyl)-1-(2-hydroxy-3-methylbutyl)-5-**

**methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) m/z:

571 (MH<sup>+</sup>). Calc'd exact mass for C<sub>32</sub>H<sub>31</sub>FN<sub>4</sub>O<sub>5</sub>: 570. <sup>1</sup>H NMR (400 MHz, CHLOROFORM-  
 d) 0.73 (d, J=6.85 Hz, 3 H) 0.81 (d, J=6.85 Hz, 3 H) 1.43 - 1.65 (m, 1 H) 2.83 (s, 3 H) 3.39 -  
 3.45 (m, 1 H) 3.94 (s, 3 H) 7.05 (d, J=7.04 Hz, 2 H) 7.20 - 7.28 (m, 3 H) 7.34 (d, J=2.35 Hz, 1  
 H) 7.38 - 7.46 (m, 3 H) 7.97 (dd, J=12.42, 1.66 Hz, 1 H) 8.31 (d, J=9.19 Hz, 1 H) 8.47 (d,  
 5 J=5.28 Hz, 1 H) 10.93 (s, 1 H)

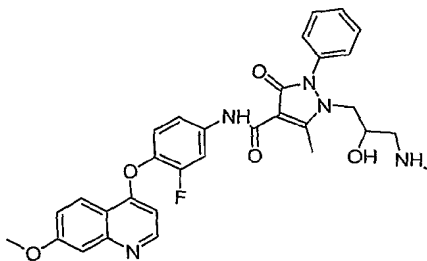
### Example 62



**N-(3-fluoro-4-(7-methoxyquinolin-4-yloxy)phenyl)-5-methyl-1-((3-methyl-2-oxooxazolidin-5-yl)methyl)-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS

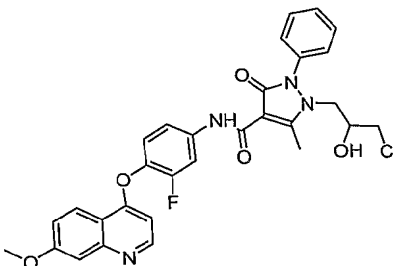
10 (ESI pos. ion) m/z: 598 (MH<sup>+</sup>). Calc'd exact mass for C<sub>32</sub>H<sub>28</sub>FN<sub>5</sub>O<sub>6</sub>: 597. <sup>1</sup>H NMR (400  
 MHz, CHLOROFORM-*d*) 10.77 (1 H, s), 8.61 (1 H, d, J=5.7 Hz), 8.28 (1 H, d, J=9.2 Hz), 7.91  
 (1 H, dd, J=12.3, 2.2 Hz), 7.68 (1 H, d, J=2.0 Hz), 7.58 (2 H, t, J=7.6 Hz), 7.48 (1 H, t, J=7.5  
 Hz), 7.38 (2 H, d, J=7.4 Hz), 7.33 (1 H, d, J=8.8 Hz), 7.23 - 7.30 (1 H, m), 7.17 (1 H, t, J=8.6  
 Hz), 6.49 (1 H, d, J=5.5 Hz), 4.42 - 4.55 (1 H, m), 4.14 (1 H, dd, J=15.7, 8.4 Hz), 4.04 (3 H, s),  
 15 3.91 (1 H, dd, J=15.7, 3.3 Hz), 3.52 (1 H, t, J=8.9 Hz), 3.06 (1 H, dd, J=9.2, 5.9 Hz), 2.77 -  
 2.93 (6 H, m), 2.67 (4 H, s)

### Example 63

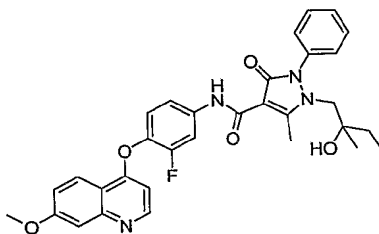


20 **N-(3-fluoro-4-(7-methoxyquinolin-4-yloxy)phenyl)-1-(2-hydroxy-3-(methylamino)propyl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)  
 m/z: 609 (MH<sup>+</sup>). Calc'd exact mass for C<sub>31</sub>H<sub>30</sub>FN<sub>5</sub>O<sub>5</sub>: 608. <sup>1</sup>H NMR (400 MHz, DMSO-*d*<sub>6</sub>)  
 11.00 (1 H, s), 8.62 (1 H, d, J=5.3 Hz), 8.23 (1 H, d, J=9.2 Hz), 7.97 (1 H, s), 7.59 (2 H, t,

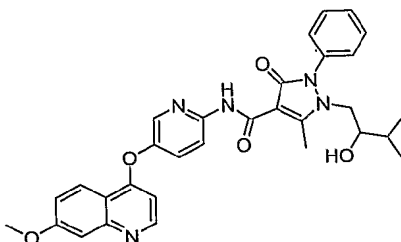
$J=7.5$  Hz), 7.26 - 7.54 (7 H, m), 6.49 (1 H, d,  $J=5.1$  Hz), 5.15 (1 H, s), 3.79 - 4.00 (5 H, m), 3.35 - 3.59 (3 H, m), 2.75 (3 H, s), 2.29 (2 H, d,  $J=5.5$  Hz), 2.07 (3 H, s), 0.99 - 1.16 (1 H, m)

**Example 64**

- 5 **1-(3-chloro-2-hydroxypropyl)-N-(3-fluoro-4-(7-methoxyquinolin-4-yloxy)phenyl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)  $m/z$ : 577 (MH<sup>+</sup>). Calc'd exact mass for C<sub>30</sub>H<sub>26</sub>ClFN<sub>4</sub>O<sub>5</sub>: 576

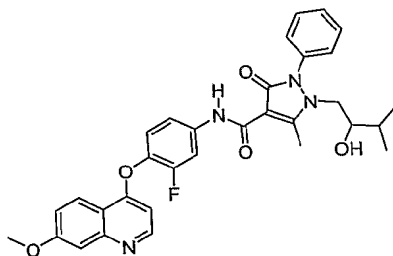
**Example 65**

- 10 **N-(3-fluoro-4-(7-methoxyquinolin-4-yloxy)phenyl)-1-(2-hydroxy-2-methylbutyl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)  $m/z$ : 571 (MH<sup>+</sup>). Calc'd exact mass for C<sub>32</sub>H<sub>31</sub>FN<sub>4</sub>O<sub>5</sub>: 570. <sup>1</sup>H NMR (400 MHz, DMSO-*d*<sub>6</sub>) 10.90 (1 H, s), 8.63 (1 H, d,  $J=5.1$  Hz), 8.24 (1 H, d,  $J=9.0$  Hz), 7.99 (1 H, dd,  $J=13.0, 2.2$  Hz), 7.64 (1 H, s), 7.59 (2 H, t,  $J=7.5$  Hz), 7.51 (1 H, t,  $J=7.4$  Hz), 7.41 - 7.47 (4 H, m), 7.36 - 7.41 (1 H, s), 7.32 (1 H, dd,  $J=9.1, 2.4$  Hz), 6.50 (1 H, d,  $J=5.3$  Hz), 4.55 - 4.65 (1 H, m), 4.30 (1 H, dd,  $J=16.0, 9.2$  Hz), 4.02 (1 H, dd,  $J=15.9, 2.6$  Hz), 3.95 (3 H, s), 3.43 (1 H, t,  $J=9.1$  Hz), 3.07 (1 H, dd,  $J=9.4, 5.5$  Hz), 2.76 (3 H, s)
- 15

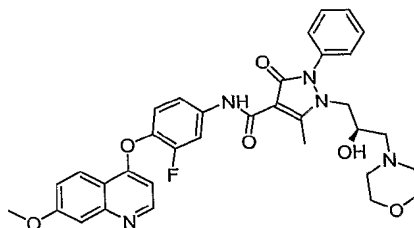
**Example 66**

- 20 **1-(2-hydroxy-3-methylbutyl)-N-(5-(7-methoxyquinolin-4-yloxy)pyridin-2-yl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)  $m/z$ : 554

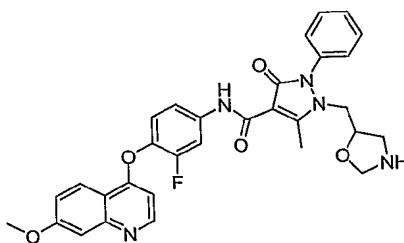
(MH<sup>+</sup>). Calc'd exact mass for C<sub>31</sub>H<sub>31</sub>N<sub>5</sub>O<sub>5</sub>: 553. <sup>1</sup>H NMR (400 MHz, DMSO-*d*<sub>6</sub>) 11.30 (1 H, s), 8.63 (1 H, d, *J*=5.3 Hz), 8.30 - 8.42 (2 H, m), 8.22 (1 H, d, *J*=9.2 Hz), 7.81 (1 H, dd, *J*=9.1, 2.8 Hz), 7.60 (2 H, t, *J*=7.5 Hz), 7.51 (1 H, t, *J*=7.3 Hz), 7.39 - 7.47 (3 H, m), 7.30 (1 H, dd, *J*=9.2, 2.5 Hz), 6.54 (1 H, d, *J*=5.3 Hz), 5.11 (1 H, d, *J*=5.7 Hz), 3.86 - 3.98 (4 H, m), 3.66 - 3.75 (1 H, m), 3.09 - 3.20 (1 H, m), 2.76 (3 H, s), 1.34 - 1.45 (1 H, m), 0.64 (3 H, d, *J*=6.7 Hz), 0.58 (3 H, d, *J*=6.7 Hz)

**Example 67**

**N-(3-fluoro-4-(7-methoxyquinolin-4-yloxy)phenyl)-1-(2-hydroxy-3-methylbutyl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) *m/z*: 571 (MH<sup>+</sup>). Calc'd exact mass for C<sub>32</sub>H<sub>31</sub>FN<sub>4</sub>O<sub>5</sub>: 570.

**Example 68**

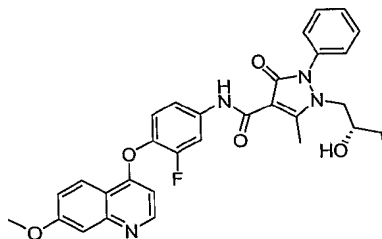
**N-(3-fluoro-4-(7-methoxyquinolin-4-yloxy)phenyl)-1-(2-hydroxy-3-morpholinopropyl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) *m/z*: 628 (MH<sup>+</sup>). Calc'd exact mass for C<sub>34</sub>H<sub>34</sub>FN<sub>5</sub>O<sub>6</sub>: 627.

**Example 69**

**N-(3-fluoro-4-(7-methoxyquinolin-4-yloxy)phenyl)-5-methyl-1-(oxazolidin-5-ylmethyl)-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) *m/z*: 584

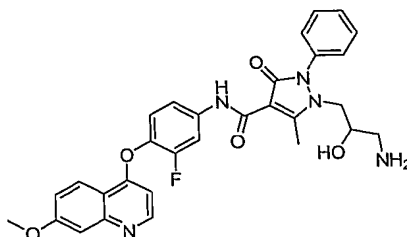
(MH<sup>+</sup>). Calc'd exact mass for C<sub>31</sub>H<sub>26</sub>FN<sub>5</sub>O<sub>6</sub>: 583. <sup>1</sup>H NMR (400 MHz, DMSO-*d*<sub>6</sub>) 11.30 (1 H, s), 8.63 (1 H, d, *J*=5.3 Hz), 8.30 - 8.42 (2 H, m), 8.22 (1 H, d, *J*=9.2 Hz), 7.81 (1 H, dd, *J*=9.1, 2.8 Hz), 7.60 (2 H, t, *J*=7.5 Hz), 7.51 (1 H, t, *J*=7.3 Hz), 7.39 - 7.47 (3 H, m), 7.30 (1 H, dd, *J*=9.2, 2.5 Hz), 6.54 (1 H, d, *J*=5.3 Hz), 5.11 (1 H, d, *J*=5.7 Hz), 3.86 - 3.98 (4 H, m), 3.66 - 3.75 (1 H, m), 3.09 - 3.20 (1 H, m), 2.76 (3 H, s), 1.34 - 1.45 (1 H, m), 0.64 (3 H, d, *J*=6.7 Hz), 0.58 (3 H, d, *J*=6.7 Hz)

### Example 70



(S)-N-(3-fluoro-4-(7-methoxyquinolin-4-yloxy)phenyl)-1-(2-hydroxybutyl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide: MS (ESI pos. ion) *m/z*: 557 (MH<sup>+</sup>). Calc'd exact mass for C<sub>31</sub>H<sub>29</sub>FN<sub>4</sub>O<sub>5</sub>: 556.

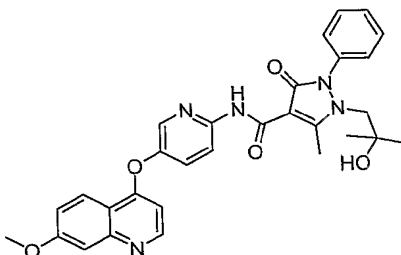
## Example 71



**1-(3-amino-2-hydroxypropyl)-N-(3-fluoro-4-(7-methoxyquinolin-4-yloxy)phenyl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) m/z:

558 (MH<sup>+</sup>). Calc'd exact mass for C<sub>30</sub>H<sub>28</sub>FN<sub>5</sub>O<sub>5</sub>: 557. <sup>1</sup>H NMR (400 MHz, DMSO-*d*<sub>6</sub>) 10.93 (1 H, s), 8.55 (1 H, d, *J*=5.3 Hz), 8.16 (1 H, d, *J*=9.0 Hz), 7.92 (1 H, dd, *J*=12.9, 2.3 Hz), 7.51 (2 H, t, *J*=7.5 Hz), 7.43 (1 H, t, *J*=7.4 Hz), 7.32 - 7.38 (4 H, m), 7.21 - 7.31 (2 H, m), 6.42 (1 H, d, *J*=5.1 Hz), 5.05 (1 H, s), 3.81 - 3.90 (4 H, m), 3.69 - 3.77 (1 H, m), 2.67 - 2.72 (3 H, m), 2.23 - 2.31 (2 H, m)

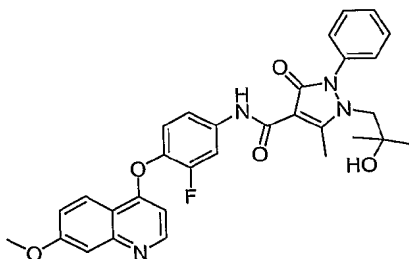
## Example 72



**1-(2-hydroxy-2-methylpropyl)-N-(5-(7-methoxyquinolin-4-yloxy)pyridin-2-yl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) m/z: 540

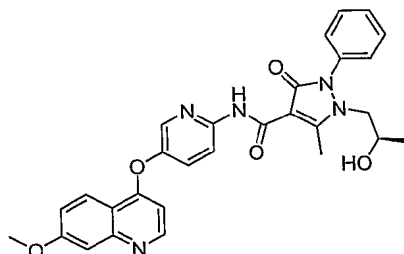
(MH<sup>+</sup>). Calc'd exact mass for C<sub>30</sub>H<sub>29</sub>N<sub>5</sub>O<sub>5</sub>: 539. <sup>1</sup>H NMR (400 MHz, CHLOROFORM-*d*) 11.23 (1 H, s), 8.58 (1 H, d, *J*=5.1 Hz), 8.37 (1 H, d, *J*=8.8 Hz), 8.20 - 8.27 (2 H, m), 7.50 (3 H, d, *J*=7.8 Hz), 7.38 - 7.44 (2 H, m), 7.26 - 7.34 (2 H, m), 7.23 (1 H, d, *J*=8.8 Hz), 6.43 (1 H, d, *J*=4.9 Hz), 3.97 (3 H, s), 3.88 (2 H, s), 2.88 (3 H, s), 2.51 (1 H, s), 1.14 (6 H, s)

## Example 73



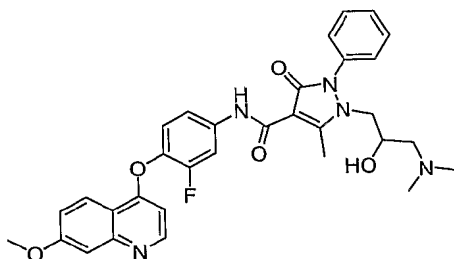
**N-(3-fluoro-4-(7-methoxyquinolin-4-yloxy)phenyl)-1-(2-hydroxy-2-methylpropyl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) m/z: 557 (MH<sup>+</sup>). Calc'd exact mass for C<sub>31</sub>H<sub>29</sub>N<sub>4</sub>O<sub>5</sub>: 556. <sup>1</sup>H NMR (400 MHz, CHLOROFORM-

*d*) 10.88 (1 H, s), 8.56 (1 H, d,  $J=5.3$  Hz), 8.27 (1 H, d,  $J=9.2$  Hz), 7.92 (1 H, dd,  $J=12.5$ , 2.0 Hz), 7.54 (2 H, t,  $J=7.6$  Hz), 7.38 - 7.49 (2 H, m), 7.29 (3 H, d,  $J=8.0$  Hz), 7.23 (1 H, dd,  $J=9.2$ , 2.3 Hz), 7.17 (1 H, t,  $J=8.7$  Hz), 6.42 (1 H, d,  $J=5.3$  Hz), 3.97 (3 H, s), 3.88 (2 H, s), 2.89 (3 H, s), 1.82 (1 H, s), 1.15 (6 H, s)

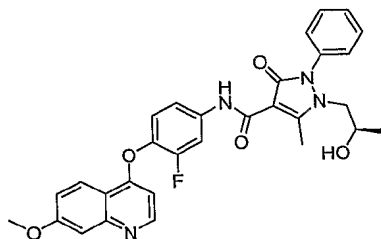
**Example 74**

**(*R*)-1-(2-hydroxypropyl)-N-(5-(7-methoxyquinolin-4-yloxy)pyridin-2-yl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)  $m/z$ : 526 (MH<sup>+</sup>). Calc'd exact mass for C<sub>29</sub>H<sub>27</sub>N<sub>5</sub>O<sub>5</sub>: 525. <sup>1</sup>H NMR (400 MHz, CHLOROFORM-*d*) 11.20 (1 H,

s), 8.55 (1 H, d,  $J=5.3$  Hz), 8.37 (1 H, d,  $J=9.0$  Hz), 8.21 - 8.28 (2 H, m), 7.46 - 7.56 (3 H, m), 7.37 - 7.45 (2 H, m), 7.21 - 7.31 (4 H, m), 6.45 (1 H, d,  $J=5.3$  Hz), 3.89 - 4.01 (4 H, m), 3.78 - 3.88 (1 H, m), 3.66 (1 H, dd,  $J=14.8$ , 2.2 Hz), 2.83 (3 H, s), 1.08 (3 H, d,  $J=6.1$  Hz)

**Example 75**

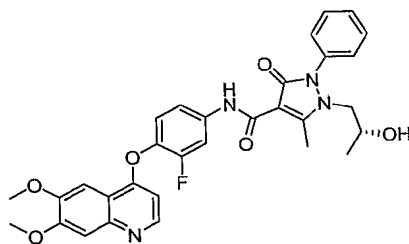
**1-(3-(dimethylamino)-2-hydroxypropyl)-N-(3-fluoro-4-(7-methoxyquinolin-4-yloxy)phenyl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)  $m/z$ : 586 (MH<sup>+</sup>). Calc'd exact mass for C<sub>32</sub>H<sub>32</sub>FN<sub>5</sub>O<sub>5</sub>: 585. <sup>1</sup>H NMR (400 MHz, CHLOROFORM-*d*) 10.88 (1 H, s), 8.58 (1 H, s), 8.23 - 8.33 (1 H, m), 7.85 - 7.97 (1 H, m), 7.11 - 7.61 (11 H, m), 6.37 - 6.46 (1 H, m), 3.97 (3 H, s), 3.79 (4 H, s), 3.48 (1 H, s), 2.87 (3 H, s), 2.11 - 2.20 (7 H, m), 1.92 - 2.02 (2 H, m).

**Example 76**

**(R)-N-(3-fluoro-4-(7-methoxyquinolin-4-yloxy)phenyl)-1-(2-hydroxypropyl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) m/z: 543

(MH<sup>+</sup>). Calc'd exact mass for C<sub>30</sub>H<sub>27</sub>FN<sub>4</sub>O<sub>5</sub>: 542. <sup>1</sup>H NMR (400 MHz, CHLOROFORM-d) 10.89 (1 H, s), 8.48 (1 H, d, J=5.3 Hz), 8.29 (1 H, d, J=9.2 Hz), 7.95 (1 H, dd, J=12.4, 1.7 Hz), 7.38 - 7.51 (1 H, m), 7.35 (1 H, d, J=2.0 Hz), 7.17 - 7.29 (3 H, m), 7.14 (2 H, d, J=7.6 Hz), 6.42 (1 H, d, J=5.3 Hz), 3.97 - 4.08 (1 H, m), 3.95 (3 H, s), 3.78 - 3.92 (2 H, m), 3.64 (1 H, d, J=12.9 Hz), 2.84 (3 H, s), 1.17 - 1.27 (4 H, m), 1.09 (3 H, d, J=5.9 Hz).

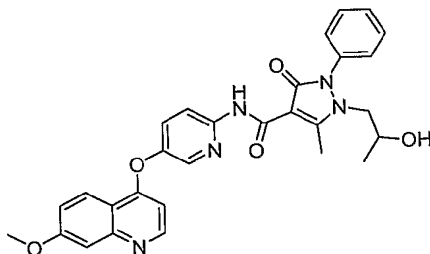
#### Example 77



**(R)-N-(4-(6,7-dimethoxyquinolin-4-yloxy)-3-fluorophenyl)-1-(2-hydroxypropyl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) m/z: 573

(MH<sup>+</sup>). Calc'd exact mass for C<sub>31</sub>H<sub>29</sub>FN<sub>4</sub>O<sub>6</sub>: 572. <sup>1</sup>H NMR (400 MHz, CHLOROFORM-d) 10.88 (1 H, s), 8.45 (1 H, d, J=5.5 Hz), 7.93 (1 H, dd, J=12.5, 2.3 Hz), 7.60 (1 H, s), 7.53 (2 H, t, J=7.5 Hz), 7.45 (1 H, t, J=7.4 Hz), 7.41 (1 H, s), 7.27 - 7.33 (3 H, m), 7.19 (1 H, t, J=8.6 Hz), 6.46 (1 H, d, J=5.3 Hz), 4.07 (3 H, s), 4.04 (3 H, s), 3.79 - 3.96 (2 H, m), 3.63 - 3.74 (1 H, m), 2.86 (3 H, s), 1.10 (3 H, d, J=6.1 Hz)

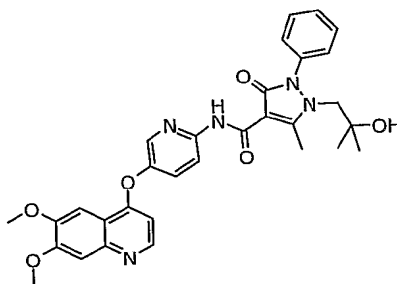
#### Example 78



**1-(2-hydroxypropyl)-N-(5-(7-methoxyquinolin-4-yloxy)pyridin-2-yl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) m/z: 526 (MH<sup>+</sup>).

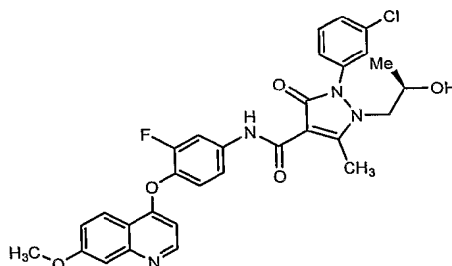
Calc'd exact mass for C<sub>29</sub>H<sub>27</sub>N<sub>5</sub>O<sub>5</sub>: 525. <sup>1</sup>H NMR (400 MHz, DMSO-d<sub>6</sub>) 11.27 (1 H, s), 8.62 (1 H, d, J=5.3 Hz), 8.36 (1 H, d, J=9.0 Hz), 8.32 (1 H, d, J=2.7 Hz), 8.22 (1 H, d, J=9.0 Hz), 7.80 (1 H, dd, J=9.0, 2.9 Hz), 7.59 (2 H, t, J=7.5 Hz), 7.51 (1 H, t, J=7.4 Hz), 7.39 - 7.45 (4 H, m), 7.30 (1 H, dd, J=9.1, 2.4 Hz), 6.54 (1 H, d, J=5.3 Hz), 5.08 (1 H, d, J=5.3 Hz), 3.94 (3 H, s), 3.88 (1 H, dd, J=15.2, 8.9 Hz), 3.55 - 3.69 (2 H, m), 2.77 (3 H, s)

## Example 79



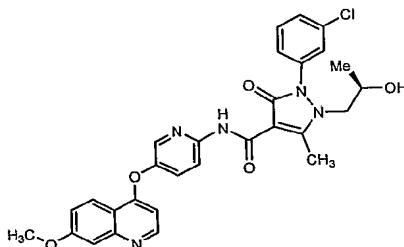
***N*-(5-(6,7-dimethoxyquinolin-4-yloxy)pyridin-2-yl)-1-(2-hydroxy-2-methylpropyl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) *m/z*: 570 (MH<sup>+</sup>). Calc'd exact mass for C<sub>31</sub>H<sub>31</sub>N<sub>5</sub>O<sub>6</sub>: 569. <sup>1</sup>H NMR (400 MHz, DMSO-*d*<sub>6</sub>) 11.25 (s, 1 H), 8.49 (d, *J*=5.2 Hz, 1 H), 8.36 (d, *J*=9.0 Hz, 1 H), 8.32 (d, *J*=2.7 Hz, 1 H), 7.79 (dd, *J*=9.0, 3.0 Hz, 1 H), 7.66 - 7.29 (m, 7 H), 6.54 (d, *J*=5.2 Hz, 1 H), 4.85 (s, 1 H), 3.95 (s, 3 H), 3.94 (s, 3 H), 3.86 (s, 2 H), 2.81 (s, 3 H), 0.96 (s, 6 H).

## Example 80

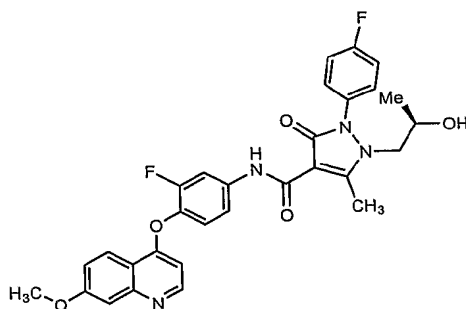


**(*R*)-2-(3-chlorophenyl)-N-(3-fluoro-4-(7-methoxyquinolin-4-yloxy)phenyl)-1-(2-hydroxypropyl)-5-methyl-3-oxo-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) *m/z*: 577 (MH<sup>+</sup>). Calc'd exact mass for C<sub>30</sub>H<sub>26</sub>ClFN<sub>4</sub>O<sub>5</sub>: 576.

## Example 81

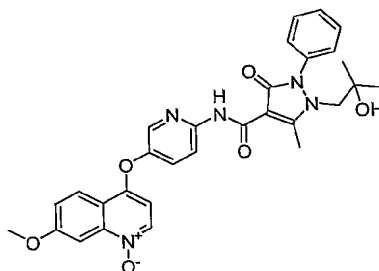


**(*R*)-2-(3-chlorophenyl)-1-(2-hydroxypropyl)-N-(5-(7-methoxyquinolin-4-yloxy)pyridin-2-yl)-5-methyl-3-oxo-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) *m/z*: 560 (MH<sup>+</sup>). Calc'd exact mass for C<sub>29</sub>H<sub>26</sub>ClN<sub>5</sub>O<sub>5</sub>: 559.

**Example 82**

**(R)-N-(3-fluoro-4-(7-methoxyquinolin-4-yloxy)phenyl)-2-(4-fluorophenyl)-1-(2-hydroxypropyl)-5-methyl-3-oxo-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos.

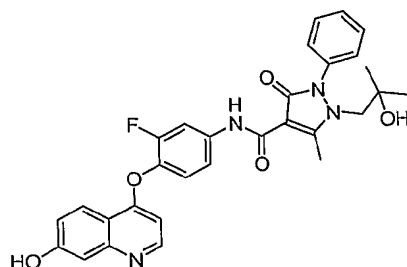
5 ion) m/z: 561 (MH<sup>+</sup>). Calc'd exact mass for C<sub>30</sub>H<sub>26</sub>F<sub>2</sub>N<sub>4</sub>O<sub>5</sub>: 560.

**Example 83**

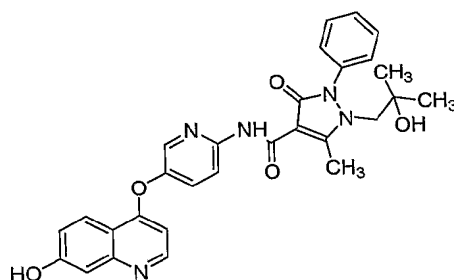
**1-(2-hydroxy-2-methylpropyl)-N-(5-(1-oxo-7-methoxyquinolin-4-yloxy)pyridin-2-yl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** A mixture of m-CPBA

10 (151 mg, 675 μmol) and 1-(2-hydroxy-2-methylpropyl)-N-(5-(7-methoxyquinolin-4-yloxy)pyridin-2-yl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide (280 mg, 519 μmol) in dichloromethane (10 mL) was stirred at room temperature for 14 h. More m-CPBA (151 mg, 675 μmol) was added, and the mixture was stirred for 24 h. The product fraction was purified from preparative HPLC to give the title compound as a white solid (25

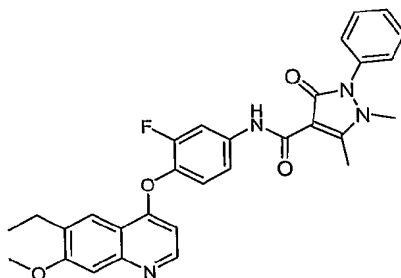
15 mg, 8.7%). Calc'd for C<sub>30</sub>H<sub>29</sub>N<sub>5</sub>O<sub>6</sub>, 555; MS (ESI pos. ion) m/z: 556 (MH<sup>+</sup>). <sup>1</sup>HNMR (400 MHz, CDCl<sub>3</sub>): 11.3 (1H, s), 8.60 (1H, d, J 6.7), 8.42 (1H, d, J 9.0), 8.32 (1H, d, J 9.4), 8.23 (1H, s), 8.00 (1H, s), 7.54-7.50 (3H, m), 7.44-7.40 (2H, m), 7.30 (2H, m), 6.52 (1H, d, J 7.1), 4.06 (3H, s), 3.88 (2H, s), 2.87 (3H, s), 1.15 (6H, s).

**Example 84**

**N-(3-Fluoro-4-(7-hydroxyquinolin-4-yloxy)phenyl)-1-(2-hydroxy-2-methylpropyl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** Calc'd for  $C_{30}H_{27}FN_4O_5$ : 542; MS (ESI pos. ion)  $m/z$ : 543 (MH<sup>+</sup>). <sup>1</sup>HNMR (400 MHz, DMSO-d<sub>6</sub>): 10.95 (1H, s, NH), 10.39 (1H, s, OH), 8.56 (1H, d, J 5.4), 8.18 (1H, d, J 7.2), 7.96 (1H, d, J 7.9), 7.58 - 7.22 (8H, m), 6.42 (1H, d, J 4.0), 5.75 (1H, s), 4.83 (1H, s, OH), 3.87 (2H, s, CH<sub>2</sub>), 2.80 (3H, s), 0.96 (6H, s).

**Example 85**

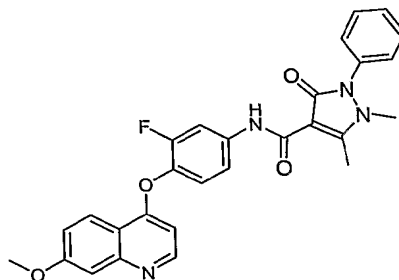
**1-(2-hydroxy-2-methylpropyl)-N-(5-(7-hydroxyquinolin-4-yloxy)pyridin-2-yl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** Calc'd for  $C_{29}H_{27}N_5O_5$ : 525; MS (ESI pos. ion)  $m/z$ : 526 (MH<sup>+</sup>).

**Example 86**

**N-(4-(6-Ethyl-7-methoxyquinolin-4-yloxy)-3-fluorophenyl)-1,5-dimethyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** Calc'd for  $C_{30}H_{27}FN_4O_4$ : 526; MS (ESI pos. ion)  $m/z$ : 527 (MH<sup>+</sup>). <sup>1</sup>HNMR (400 MHz, CDCl<sub>3</sub>): 10.88 (1H), 8.54 (1H, d, J = 5.1), 8.09 (1H, s), 7.92 (1H, dd, J 2.0, 13.8), 7.57 (2H, t, J 7.5), 7.48 (1H, t, J 7.2), 7.37 (2H, m), 7.32 (1H, d),

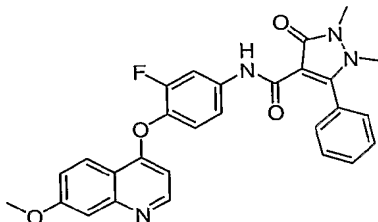
7.17 (1H, t, J 8.8), 6.39 (1H, d, 4.9), 3.99 (3H, s), 3.38 (3H, s), 2.84 (2H, m) 2.81 (3H, s), 1.32 (3H, t, J 7.5).

### Example 87



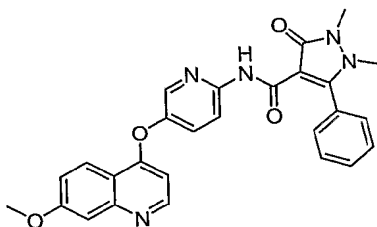
- 5 **N-(3-Fluoro-4-(7-methoxyquinolin-4-yloxy)phenyl)-1,5-dimethyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** Calc'd for  $C_{28}H_{23}FN_4O_4$ : 498; MS (ESI pos. ion) m/z: 499 (MH<sup>+</sup>). <sup>1</sup>HNMR (400 MHz, CDCl<sub>3</sub>): 10.88 (1H), 8.58 (1H, d, J = 5.7), 8.27 (1H, d, J = 9.2), 7.92 (1H, dd, J 2.1, 12.5), 7.57 (2H, t, J 7.8), 7.48 (1H, t, J 7.4), 7.41 (ds, J 2.4), 7.37 (2H, J 7.6), 7.29 (1H, d), 7.22 (1H, dd, J 2.4, 9.2), 7.17 (2H, t, J 8.6), 6.41 (1H, d, 5.3), 3.97 (3H, s), 3.38 (3H, s), 2.80 (3H, s).

### Example 88



- N-(3-fluoro-4-(7-Methoxyquinolin-4-yloxy)phenyl)-1,2-dimethyl-3-oxo-5-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** A mixture of HATU (458 mg, 1206  $\mu$ mol), 1,2-dimethyl-3-oxo-5-phenyl-2,3-dihydro-1H-pyrazole-4-carboxylic acid (140 mg, 603  $\mu$ mol), 3-fluoro-4-(7-methoxyquinolin-4-yloxy)benzenamine (258 mg, 908  $\mu$ mol), and Triethylamine (254  $\mu$ L, 1809  $\mu$ mol) in DMF (2 mL) was stirred at 60°C overnight. The mixture was diluted with EtOAc (10 mL). The mixture was transferred to a separatory funnel with EtOAc (20 mL) and was washed with NaOH (1N, 10 mL), H<sub>2</sub>O (2x10 mL), NaHCO<sub>3</sub> (sat), NaCl (sat), and dried over Na<sub>2</sub>SO<sub>4</sub>. The residue after concentration was purified on silica gel and the product was triturated with EtOAc-hexane (1:2) to afford a pink solid (106 mg, 35%). Calc'd for  $C_{28}H_{23}FN_4O_4$ : 498; MS (ESI pos. ion) m/z: 499 (MH<sup>+</sup>). <sup>1</sup>HNMR (CDCl<sub>3</sub>, 400 MHz): 11.04 (1H, s), 8.56 (1H, d, J 5.3), 7.88 (1H, d, J 14.7), 7.55-7.40 (5H, m), 7.47 (1H, s), 7.35 (1H, d), 7.22 (1H, d, J 9.1), 7.13 (1H, t, J 8.8), 6.36 (1H, d, J 5.3), 3.96 (3H, s), 3.62 (3H, s), 3.41 (3H, s).

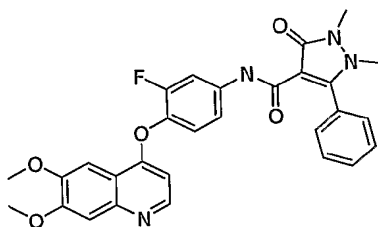
## Example 89



**N-(5-(7-Methoxyquinolin-4-yloxy)pyridin-2-yl)-1,2-dimethyl-3-oxo-5-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** Calc'd for  $C_{27}H_{23}N_5O_4$ : 481; MS (ESI pos. ion) m/z: 482

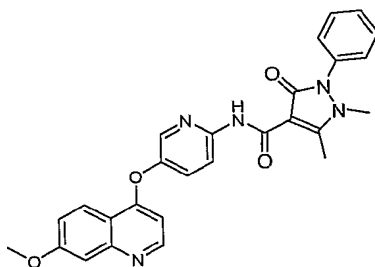
5 (MH<sup>+</sup>). <sup>1</sup>HNMR (CDCl<sub>3</sub>, 400 MHz): 11.46 (1H, s), 8.58 (1H, d, J 5.3), 8.30 (1H, d, J 9.0), 8.25-8.21 (2H, m), 7.56-7.40 (7H), 7.22 (1H, dd, J 2.3, 9.0), 6.39 (1H, d, J 5.3), 3.97 (3H, s), 3.62 (3H, s), 3.40 (3H, s).

## Example 90



10 **N-(4-(6,7-Dimethoxyquinolin-4-yloxy)-3-fluorophenyl)-1,2-dimethyl-3-oxo-5-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** Calc'd for  $C_{27}H_{23}N_5O_4$ : 528; MS (ESI pos. ion) m/z: 529(MH<sup>+</sup>). <sup>1</sup>HNMR (CDCl<sub>3</sub>, 400 MHz): 11.05 (1H, s), 8.46 (1H, d, J 5.2), 7.88 (1H, dd, J 2.2, 12.5), 7.53-7.58 (4H, m), 7.47-7.49 (2H, m), 7.40 (1H, s), 7.32 (1H, d, 8), 7.14 (1H, t, J 8.7), 6.39 (1H, d, J 5.1), 4.05 (3H, s), 4.04 (3H, s), 3.62 (3H, s), 3.41 (3H, s).

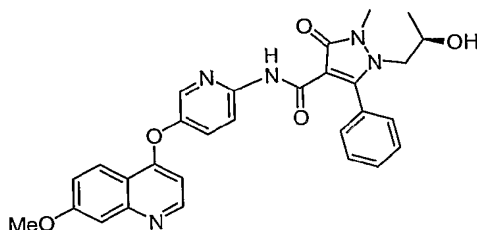
## Example 91



**N-(5-(7-Methoxyquinolin-4-yloxy)pyridin-2-yl)-1,5-dimethyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** Calc'd for  $C_{27}H_{23}N_5O_4$ : 481; MS (ESI pos. ion) m/z: 482

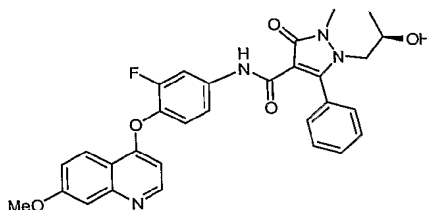
20 (MH<sup>+</sup>). <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz): 11.27 (1H, s), 8.60 (1H, d, J 5.1), 8.38 (1H, d, J 9.0), 8.23 (2H, m), 7.53 (3H, m), 7.47 (1H, m), 7.41 (1H, d), 7.37 (2H, d, J 7.4), 7.23 (1H, dd, J 2.4, 9.7), 6.42 (1H, d, J 5.1), 3.97 (3H, s), 3.37 (3H, s), 2.80 (3H, s).

## Example 92



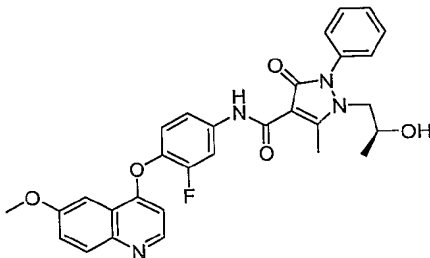
**(R)-1-(2-Hydroxypropyl)-N-(5-(7-methoxyquinolin-4-yloxy)pyridin-2-yl)-2-methyl-3-oxo-5-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** Calc'd for  $C_{29}H_{27}N_5O_5$ : 525; MS (ESI pos. ion)  $m/z$ : 526 (MH<sup>+</sup>). <sup>1</sup>HNMR (CDCl<sub>3</sub>, 400 MHz): 11.28 (1H, s), 8.53 (1H, d, J 5.1), 8.24-8.21 (3H, m), 7.49-7.39 (7H, m), 7.22 (1H, bd, J 9.2), 6.38 (1H, d, J 5.1), 4.7 (1H, b), 3.96 (4H, bs), 3.75 (2H, m), 3.63 (3H, s), 1.04 (3H, d, J 6.3).

## Example 93



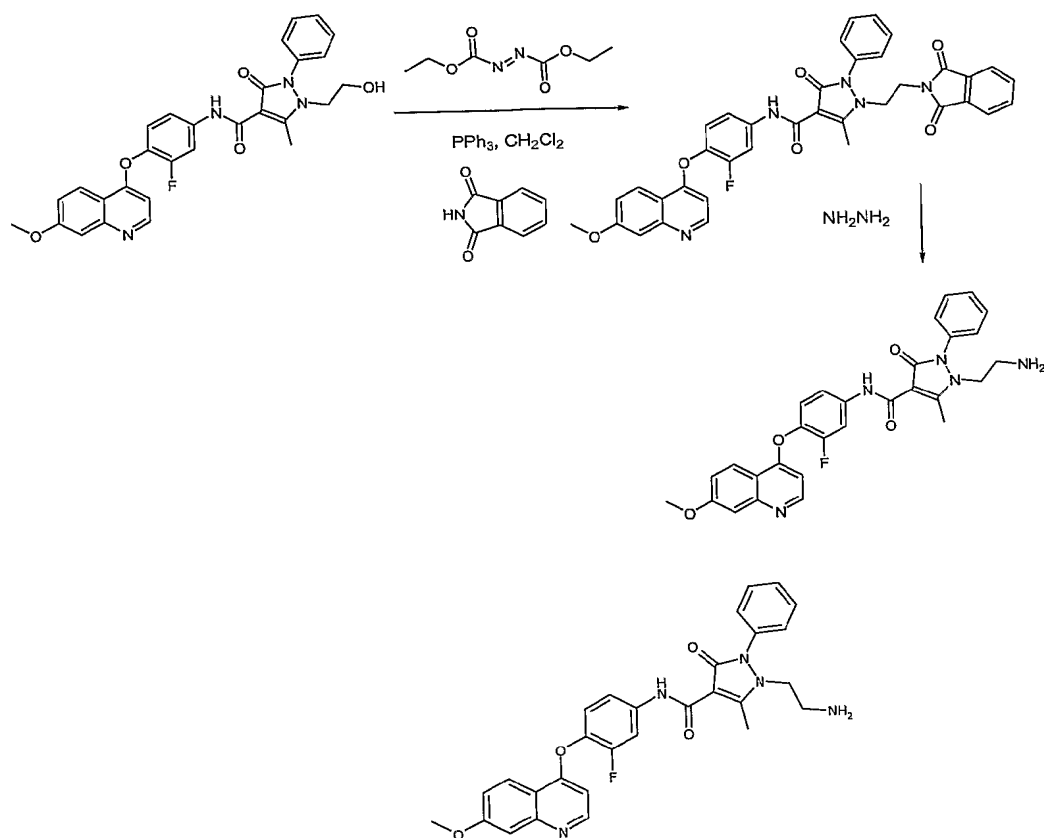
**(R)-N-(3-Fluoro-4-(7-methoxyquinolin-4-yloxy)phenyl)-1-(2-hydroxypropyl)-2-methyl-3-oxo-5-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** Calc'd for  $C_{30}H_{27}FN_4O_5$ : 542; MS (ESI pos. ion)  $m/z$ : 543 (MH<sup>+</sup>). <sup>1</sup>HNMR (CDCl<sub>3</sub>, 400 MHz): 11.05 (1H, s), 8.55 (1H, d, J 5.3), 8.26 (1H, d, J 9.2), 7.86 (1H, d, J 12.5), 7.53-7.45 (5H, m), 7.39 (1H, d, J 2.3), 7.32 (1H, d, J 9.7), 7.77 (1H, dd, J 9.2, 2.6), 7.14 (1H, t, J 8.8), 6.37 (1H, J 5.3), 3.96 (3H, s), 3.90 (1H, m), 3.76 (2H, d, J 6.0), 3.62 (3H, s), 1.94 (1H, d, J 4.3), 1.04 (3H, d, J 6.2).

## Example 94



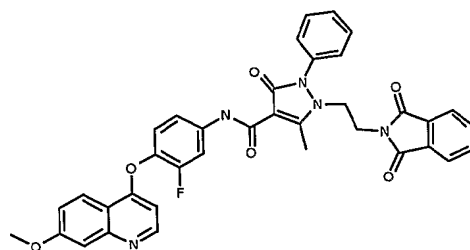
**(S)-N-(3-fluoro-4-(6-methoxyquinolin-4-yloxy)phenyl)-1-(2-hydroxypropyl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)  $m/z$ : 543 (MH<sup>+</sup>). Calc'd exact mass for  $C_{30}H_{27}FN_4O_5$ : 542. <sup>1</sup>HNMR (300 MHz, CDCl<sub>3</sub>): 1.01 (d, J=5.85 Hz, 3H), 2.72-2.79 (m, 3H), 3.55 (d, J=12.28 Hz, 1H), 3.69-3.93 (m, 3H), 6.44 (d,

J=4.53 Hz, 1H), 7.02 (d, J=6.72 Hz, 2H), 7.04 (s, 2H), 7.27-7.40 (m, 4H), 7.56 (d, J=2.78 Hz, 1H), 7.78-7.91 (m, 2H), 8.31 (d, J=5.26 Hz, 1H), 10.80 (s, 1H).

**Example 95**

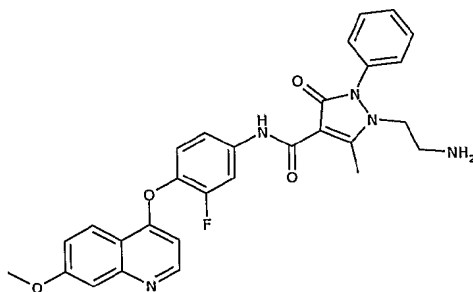
5

**1-(2-aminoethyl)-N-(3-fluoro-4-((7-(methoxy)-4-quinolinyl)oxy) phenyl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide**



- 10 **Step 1: 1-(2-(1,3-dioxo-1,3-dihydro-2H-isoindol-2-yl)ethyl)-N-(3-fluoro-4-((7-(methoxy)-4-quinolinyl)oxy)phenyl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide.** To a solution of N-(3-fluoro-4-(7-methoxyquinolin-4-yloxy)phenyl)-1-(2-hydroxyethyl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide (0.20 g, 0.38 mmol) and phthalimide (0.11 g, 0.76 mmol) in 10 mL of CH<sub>2</sub>Cl<sub>2</sub> was added triphenyl
- 15 phosphine (0.13 ml, 0.57 mmol), followed by diethyl azodicarboxylate (0.089 ml, 0.57 mmol)

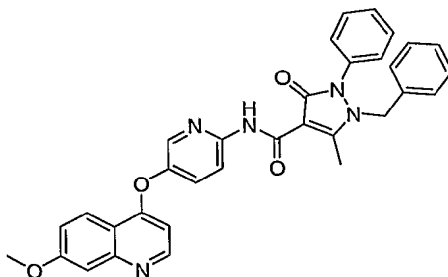
via a syringe. The reaction mixture was stirred at RT for 16 hours. The resulting solution was concentrated in vacuo, and the residue was purified by silica gel chromatography (EtOAc to 10% MeOH/EtOAc) to give the title compound as a light yellow solid (0.22 g, 88% yield). MS (ESI pos. ion)  $m/z$ : 658 (MH<sup>+</sup>). Calc'd exact mass for C<sub>37</sub>H<sub>28</sub>FN<sub>5</sub>O<sub>6</sub>: 657. <sup>1</sup>H NMR (300 MHz, MeOH) 8.53 (1 H, d,  $J$ =5.5 Hz), 8.28 (1 H, d,  $J$ =9.0 Hz), 7.80 - 7.91 (5 H, m), 7.50 (1 H, s), 7.48 (2 H, d,  $J$ =3.0 Hz), 7.26 - 7.36 (6 H, m), 6.49 (1 H, dd,  $J$ =5.4, 1.0 Hz), 4.27 (2 H, t,  $J$ =5.5 Hz), 3.97 (3 H, s), 3.74 (2 H, t,  $J$ =5.5 Hz), 2.68 (3 H, s).



**Step 2: 1-(2-aminoethyl)-N-(3-fluoro-4-((7-methoxy)-4-quinolinyl)oxy)phenyl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide.** To a solution of 1-(2-(1,3-dioxoisindolin-2-yl)ethyl)-N-(3-fluoro-4-(7-methoxyquinolin-4-yloxy)phenyl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide (0.20 g, 0.30 mmol) in 1:1 H<sub>2</sub>O/EtOH was added hydrazine (0.049 g, 1.5 mmol). The reaction was heated to 50°C for 8 hours and then cooled to RT. The reaction mixture was then diluted with 20 mL of satd. NaHCO<sub>3</sub> aq. solution and 60 mL of EtOAc. The organic phase was separated and washed with 30 mL of brine, dried over Na<sub>2</sub>SO<sub>4</sub> and concentrated in vacuo. The residue was washed with 20% hexane in EtOAc to give the title compound as a light yellow solid (0.13 g, 81% yield).

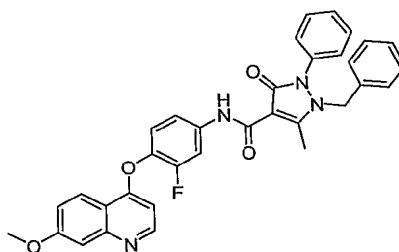
MS (ESI pos. ion)  $m/z$ : 528 (MH<sup>+</sup>). Calc'd exact mass for C<sub>29</sub>H<sub>26</sub>FN<sub>5</sub>O<sub>4</sub>: 527. <sup>1</sup>H NMR (300 MHz, MeOH) 8.54 (1 H, d,  $J$ =5.5 Hz), 8.30 (1 H, d,  $J$ =9.2 Hz), 7.93 - 7.97 (1 H, m), 7.54 - 7.66 (3 H, m), 7.44 - 7.51 (2 H, m), 7.27 - 7.38 (4 H, m), 6.50 (1 H, dd,  $J$ =5.5, 0.9 Hz), 3.92 - 4.02 (5 H, m), 2.83 (3 H, s), 2.68 (2 H, t,  $J$ =6.7 Hz).

#### Example 96

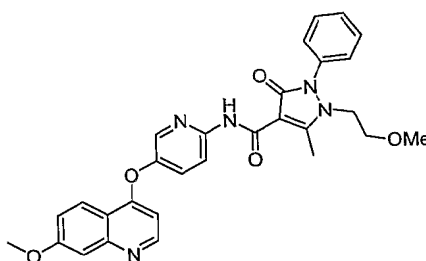


**5-methyl-N-(5-((7-(methyloxy)-4-quinolinyl)oxy)-2-pyridinyl)-3-oxo-2-phenyl-1-****(phenylmethyl)-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)  $m/z$ : 558(MH<sup>+</sup>). Calc'd exact mass for C<sub>33</sub>H<sub>27</sub>N<sub>5</sub>O<sub>4</sub>: 557. 1H NMR (300 MHz, *CHLOROFORM-d*)11.40 (1 H, s), 8.80 (1 H, d,  $J=6.4$  Hz), 8.49 (1 H, d,  $J=9.2$  Hz), 8.39 (1 H, d,  $J=9.4$  Hz), 8.27(1 H, d,  $J=2.6$  Hz), 7.95 (1 H, d,  $J=2.1$  Hz), 7.41 - 7.58 (5 H, m), 7.31 (1 H, d,  $J=2.3$  Hz), 7.23- 7.30 (4 H, m), 6.86 - 6.91 (1 H, m), 6.85 (1 H, s), 6.73 (1 H, d,  $J=6.6$  Hz), 4.98 (2 H, s), 4.07

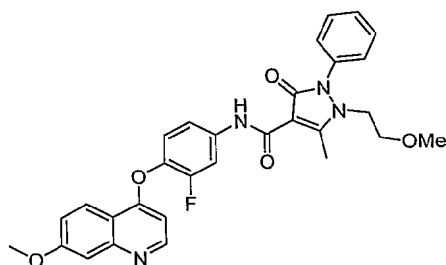
(3 H, s), 2.84 (3 H, s).

**Example 97****1-benzyl-N-(3-fluoro-4-(7-methoxyquinolin-4-yloxy)phenyl)-5-methyl-3-oxo-2-phenyl-2,3-****dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)  $m/z$ : 575 (MH<sup>+</sup>). Calc'd exactmass for C<sub>34</sub>H<sub>27</sub>FN<sub>4</sub>O<sub>4</sub>: 574. 1H NMR (300 MHz, *CHLOROFORM-d*) 11.02 (1 H, s), 8.78 (1H, s), 8.41 (1 H, d,  $J=9.4$  Hz), 8.03 (1 H, dd,  $J=12.5, 1.8$  Hz), 7.89 (1 H, s), 7.20 - 7.54 (11 H,

m), 6.80 - 6.91 (2 H, m), 6.75 (1 H, s), 4.99 (2 H, s), 4.06 (3 H, s), 2.85 (3 H, s).

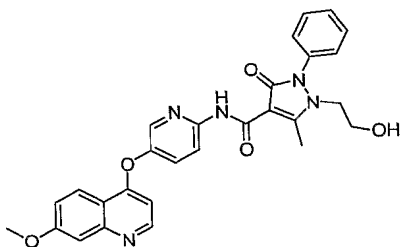
**Example 98****5-methyl-1-(2-(methyloxy)ethyl)-N-(5-((7-(methyloxy)-4-quinolinyl)oxy)-2-pyridinyl)-3-****oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)  $m/z$ : 526(MH<sup>+</sup>). Calc'd exact mass for C<sub>29</sub>H<sub>27</sub>N<sub>5</sub>O<sub>5</sub>: 525. 1H NMR (300 MHz, *MeOH*) 8.81 (1 H, d, $J=6.8$  Hz), 8.46 - 8.59 (2 H, m), 8.34 (1 H, d,  $J=2.4$  Hz), 7.84 (1 H, dd,  $J=9.2, 2.8$  Hz), 7.43 -7.64 (7 H, m), 7.00 (1 H, d,  $J=6.8$  Hz), 4.06 - 4.15 (5 H, m), 3.25 - 3.40 (5 H, m), 2.82 (3 H, s).

## Example 99



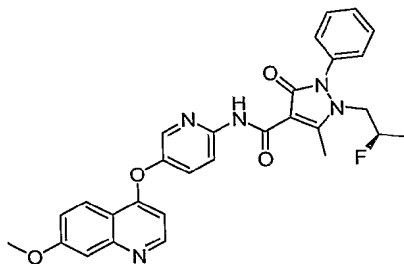
**N-(3-fluoro-4-((7-(methoxy)-4-quinolinyl)oxy)phenyl)-5-methyl-1-(2-(methoxy)ethyl)-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)  $m/z$ : 543 (MH<sup>+</sup>). Calc'd exact mass for C<sub>30</sub>H<sub>27</sub>FN<sub>4</sub>O<sub>5</sub>: 542. <sup>1</sup>H NMR (300 MHz, *CHLOROFORM-d*) 10.90 (1 H, s), 8.59 (1 H, d,  $J=5.3$  Hz), 8.27 (1 H, d,  $J=9.2$  Hz), 7.93 (1 H, dd,  $J=12.5, 2.4$  Hz), 7.52 - 7.59 (1 H, m), 7.14 - 7.48 (8 H, m), 6.41 (1 H, dd,  $J=5.3, 0.9$  Hz), 3.94 - 4.02 (5 H, m), 3.34 (2 H, t,  $J=5.0$  Hz), 3.25 (3 H, s), 2.83 (3 H, s).

## Example 100



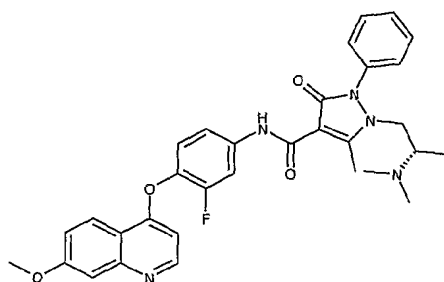
**1-(2-hydroxyethyl)-5-methyl-N-(5-((7-(methoxy)-4-quinolinyl)oxy)-2-pyridinyl)-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)  $m/z$ : 512 (MH<sup>+</sup>). Calc'd exact mass for C<sub>28</sub>H<sub>25</sub>N<sub>5</sub>O<sub>5</sub>: 511. <sup>1</sup>H NMR (300 MHz, *MeOH*) 8.81 (1 H, d,  $J=6.8$  Hz), 8.44 - 8.58 (2 H, m), 8.33 (1 H, s), 7.83 (1 H, d,  $J=9.0$  Hz), 7.43 - 7.64 (7 H, m), 7.00 (1 H, d,  $J=6.6$  Hz), 4.08 (3 H, s), 4.03 (2 H, t,  $J=4.4$  Hz), 3.51 (2 H, t,  $J=4.3$  Hz), 2.84 (3 H, s).

## Example 101



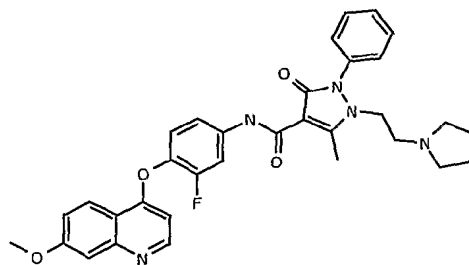
**1-((2R)-2-fluoropropyl)-5-methyl-N-(5-((7-(methoxy)-4-quinolinyl)oxy)-2-pyridinyl)-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)  $m/z$ : 528 (MH<sup>+</sup>). Calc'd exact mass for C<sub>29</sub>H<sub>26</sub>FN<sub>5</sub>O<sub>4</sub>: 527.

## Example 102



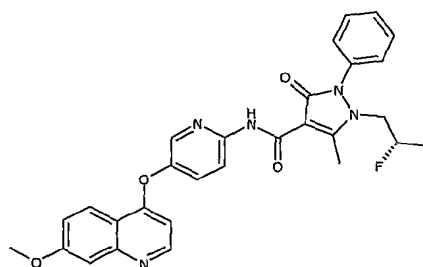
**(S)-1-(2-((dimethylamino)propyl)-N-(3-fluoro-4-((7-methoxyquinolin-4-yl)oxy)phenyl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)  $m/z$ : 570 (MH<sup>+</sup>). Calc'd exact mass for C<sub>32</sub>H<sub>32</sub>FN<sub>5</sub>O<sub>4</sub>: 569. <sup>1</sup>H NMR (300 MHz, MeOH) 8.82 (1 H, d,  $J=6.6$  Hz), 8.53 (1 H, d,  $J=9.2$  Hz), 8.01 (1 H, dd,  $J=12.8, 2.3$  Hz), 7.35 - 7.68 (9 H, m), 6.99 (1 H, dd,  $J=6.8, 0.9$  Hz), 4.27 - 4.45 (2 H, m), 4.08 (3 H, s), 3.42 - 3.54 (1 H, m), 2.87 (3 H, s), 2.67 (6 H, s), 1.23 (3 H, d,  $J=6.8$  Hz).

## Example 103



**N-(3-fluoro-4-((7-(methoxy)-4-quinolinyl)oxy)phenyl)-5-methyl-3-oxo-2-phenyl-1-(2-(1-pyrrolidinyl)ethyl)-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)  $m/z$ : 582 (MH<sup>+</sup>). Calc'd exact mass for C<sub>33</sub>H<sub>32</sub>FN<sub>5</sub>O<sub>4</sub>: 581. <sup>1</sup>H NMR (300 MHz, MeOH) 8.82 (1 H, d,  $J=6.8$  Hz), 8.55 (1 H, d,  $J=9.2$  Hz), 8.02 (1 H, dd,  $J=12.7, 1.8$  Hz), 7.37 - 7.68 (9 H, m), 7.00 (1 H, d,  $J=6.6$  Hz), 4.27 - 4.38 (2 H, m), 4.09 (3 H, s), 3.21 - 3.34 (6 H, m), 2.87 (3 H, s), 2.01 (4H, m).

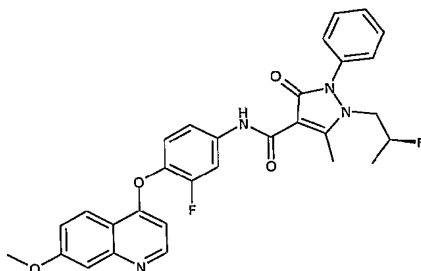
## Example 104



**1-((2S)-2-fluoropropyl)-5-methyl-N-(5-((7-(methoxy)-4-quinolinyl)oxy)-2-pyridinyl)-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)  $m/z$ : 528

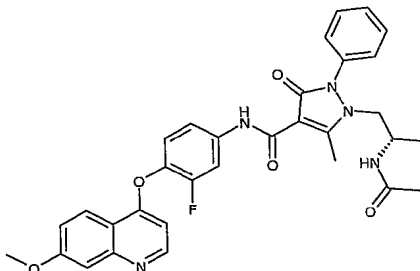
(MH<sup>+</sup>). Calc'd exact mass for C<sub>29</sub>H<sub>26</sub>FN<sub>5</sub>O<sub>4</sub>: 527. <sup>1</sup>H NMR (300 MHz, *MeOH*) 8.81 (1 H, d, *J*=6.8 Hz), 8.44 - 8.58 (2 H, m), 8.34 (1 H, d, *J*=2.6 Hz), 7.83 (1 H, dd, *J*=9.1, 2.9 Hz), 7.41 - 7.64 (7 H, m), 7.00 (1 H, d, *J*=6.8 Hz), 4.09 (3H, s), 3.88 - 4.12 (1 H, m), 3.35 (2 H, m), 2.77 - 2.85 (3 H, m), 1.13 - 1.33 (3 H, m).

5

**Example 105**

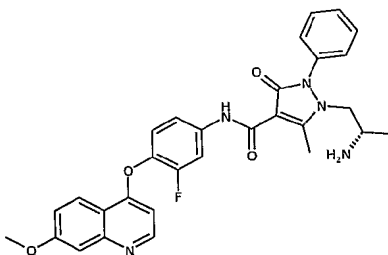
**N-(3-fluoro-4-((7-(methoxy)-4-quinolinyl)oxy)phenyl)-1-((2S)-2-fluoropropyl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) *m/z*: 545 (MH<sup>+</sup>). Calc'd exact mass for C<sub>30</sub>H<sub>26</sub>F<sub>2</sub>N<sub>4</sub>O<sub>4</sub>: 544.

10

**Example 106**

**1-((2S)-2-(acetilamino)propyl)-N-(3-fluoro-4-((7-(methoxy)-4-quinolinyl)oxy)phenyl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) *m/z*: 584 (MH<sup>+</sup>). Calc'd exact mass for C<sub>32</sub>H<sub>30</sub>FN<sub>5</sub>O<sub>5</sub>: 583. <sup>1</sup>H NMR (300 MHz, *MeOH*) 8.83 (1 H, d, *J*=6.6 Hz), 8.57 (1 H, d, *J*=9.4 Hz), 8.04 (1 H, dd, *J*=12.9, 1.8 Hz), 7.39 - 7.66 (9 H, m), 7.00 (1 H, dd, *J*=6.8, 1.1 Hz), 4.10 (3H, s), 3.83 - 4.11 (3 H, m), 2.80 (3 H, s), 1.84 (3 H, s), 0.98 (3 H, d, *J*=6.4 Hz).

15

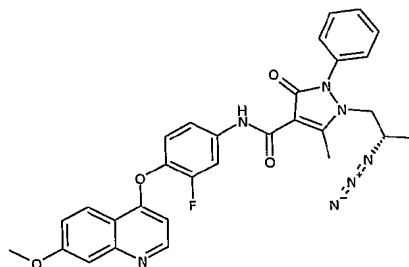
**Example 107**

20

**1-((2S)-2-aminopropyl)-N-(3-fluoro-4-((7-(methoxy)-4-quinolinyl)oxy)phenyl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) *m/z*: 542

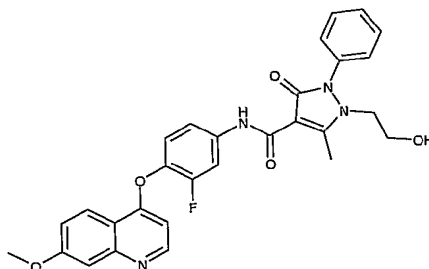
(MH<sup>+</sup>). Calc'd exact mass for C<sub>30</sub>H<sub>28</sub>FN<sub>5</sub>O<sub>4</sub>: 541. <sup>1</sup>H NMR (300 MHz, *MeOH*) 8.83 (1 H, d, *J*=6.8 Hz), 8.55 (1 H, d, *J*=9.2 Hz), 8.04 (1 H, dd, *J*=12.8, 1.7 Hz), 7.39 - 7.69 (9 H, m), 6.99 (1 H, dd, *J*=6.6, 0.9 Hz), 4.06 - 4.31 (2 H, m), 4.09 (3H, s), 3.28 - 3.41 (1 H, m), 2.86 (3 H, s), 1.14 (3 H, d, *J*=6.6 Hz).

5

**Example 108**

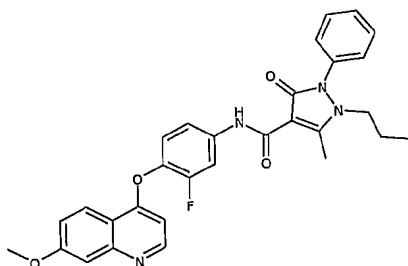
**1-((2S)-2-azidopropyl)-N-(3-fluoro-4-((7-(methyloxy)-4-quinolinyl)oxy)phenyl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) *m/z*: 568 (MH<sup>+</sup>). Calc'd exact mass for C<sub>30</sub>H<sub>26</sub>FN<sub>7</sub>O<sub>4</sub>: 567. <sup>1</sup>H NMR (300 MHz, *CHLOROFORM-d*) 10.84 (1 H, s), 8.59 (1 H, d, *J*=5.3 Hz), 8.27 (1 H, d, *J*=9.2 Hz), 7.92 (1 H, dd, *J*=12.6, 2.4 Hz), 7.53 - 7.60 (2 H, m), 7.14 - 7.50 (7 H, m), 6.41 (1 H, dd, *J*=5.3, 0.9 Hz), 3.96 (3 H, s), 3.56 - 3.84 (3 H, m), 2.85 (3 H, s), 1.15 (3 H, d, *J*=6.6 Hz).

10

**Example 109**

**N-(3-fluoro-4-((7-(methyloxy)-4-quinolinyl)oxy)phenyl)-1-(2-hydroxyethyl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) *m/z*: 529 (MH<sup>+</sup>). Calc'd exact mass for C<sub>29</sub>H<sub>25</sub>FN<sub>4</sub>O<sub>5</sub>: 528.

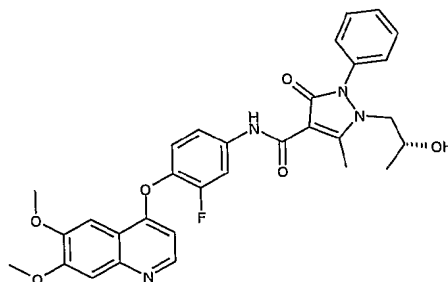
15

**Example 110**

**N-(3-fluoro-4-((7-(methyloxy)-4-quinolinyl)oxy)phenyl)-5-methyl-3-oxo-2-phenyl-1-propyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) *m/z*: 527 (MH<sup>+</sup>).

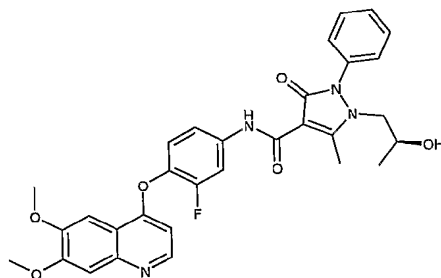
20

Calc'd exact mass for  $C_{30}H_{27}FN_4O_4$ : 526.  $^1H$  NMR (400 MHz, *CHLOROFORM-d*) 10.91 (1 H, s), 8.59 (1 H, d,  $J=5.3$  Hz), 8.27 (1 H, d,  $J=9.0$  Hz), 7.91 - 7.93 (2 H, d,  $J=2.0$  Hz), 7.41 - 7.58 (4 H, m), 7.14 - 7.38 (4 H, m), 6.42 (1 H, d,  $J=5.3$  Hz), 3.96 (3 H, s), 3.76 (2 H, t,  $J=7.2$  Hz), 2.80 (3 H, s), 1.45 - 1.56 (2 H, m,  $J=7.4, 7.4, 7.4, 7.4, 7.4$  Hz), 0.80 (3 H, t,  $J=7.4$  Hz).

**Example 111**

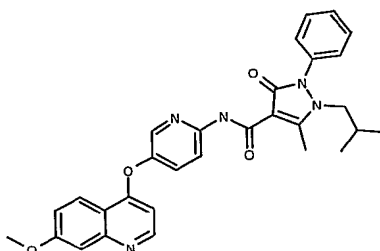
**N-(4-((6,7-bis(methoxy)-4-quinolinyl)oxy)-3-fluorophenyl)-1-((2R)-2-hydroxypropyl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) m/z:

573 (MH<sup>+</sup>). Calc'd exact mass for  $C_{31}H_{29}FN_4O_6$ : 572.

**Example 112**

**N-(4-((6,7-bis(methoxy)-4-quinolinyl)oxy)-3-fluorophenyl)-1-((2S)-2-hydroxypropyl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) m/z:

573 (MH<sup>+</sup>). Calc'd exact mass for  $C_{31}H_{29}FN_4O_6$ : 572.

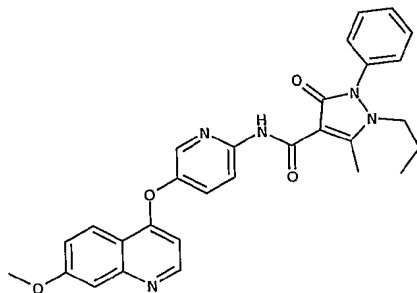
**Example 113**

**5-methyl-N-(5-((7-(methoxy)-4-quinolinyl)oxy)-2-pyridinyl)-1-(2-methylpropyl)-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) m/z: 524 (MH<sup>+</sup>).

Calc'd exact mass for  $C_{30}H_{29}N_5O_4$ : 523.  $^1H$  NMR (400 MHz, *CHLOROFORM-d*) 11.28 (1 H,

s), 8.60 (1 H, d,  $J=5.3$  Hz), 8.37 (1 H, d,  $J=8.8$  Hz), 8.24 (1 H, d,  $J=6.1$  Hz), 8.23 (1 H, s), 7.41 - 7.56 (5 H, m), 7.35 (2 H, d,  $J=8.0$  Hz), 7.23 (1 H, dd,  $J=9.2, 2.3$  Hz), 6.43 (1 H, d,  $J=5.3$  Hz), 3.97 (3 H, s), 3.64 (2 H, d,  $J=7.4$  Hz), 2.81 (3 H, s), 1.84 (1 H, dt,  $J=13.8, 6.9$  Hz), 0.77 (6 H, d,  $J=6.7$  Hz).

5

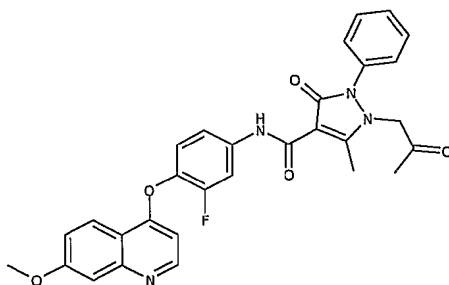
**Example 114**

**5-methyl-N-(5-((7-(methyloxy)-4-quinolinyl)oxy)-2-pyridinyl)-3-oxo-2-phenyl-1-propyl-**

**2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)  $m/z$ : 510 (MH<sup>+</sup>). Calc'd exact mass for C<sub>29</sub>H<sub>27</sub>N<sub>5</sub>O<sub>4</sub>: 509. <sup>1</sup>H NMR (400 MHz, CHLOROFORM-*d*) 11.29 (1 H, s), 8.61 (1 H, d,  $J=5.5$  Hz), 8.38 (1 H, d,  $J=9.0$  Hz), 8.22 - 8.27 (2 H, m), 7.32 - 7.57 (7 H, m), 7.22 - 7.30 (1 H, m), 6.46 (1 H, d,  $J=5.5$  Hz), 3.98 (3 H, s), 3.76 (2 H, t,  $J=7.2$  Hz), 2.79 - 2.83 (3 H, m), 1.42 - 1.56 (2 H, m,  $J=7.4, 7.4, 7.4, 7.4$  Hz), 0.70 - 0.86 (3 H, t,  $J=7.4$  Hz).

10

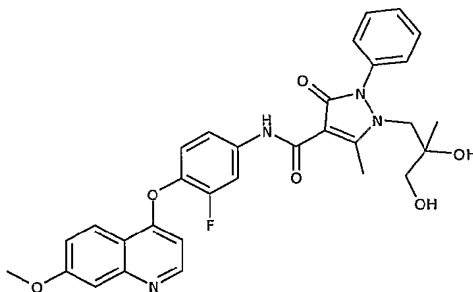
## Example 115



**N-(3-fluoro-4-((7-(methoxy)-4-quinolinyl)oxy)phenyl)-5-methyl-3-oxo-1-(2-oxopropyl)-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) m/z: 541 (MH<sup>+</sup>).

- 5 Calc'd exact mass for C<sub>30</sub>H<sub>25</sub>FN<sub>4</sub>O<sub>5</sub>: 540. <sup>1</sup>H NMR (300 MHz, *CHLOROFORM-d*) 10.86 (1 H, s), 8.60 (1 H, d, *J*=5.3 Hz), 8.28 (1 H, d, *J*=9.2 Hz), 7.92 (1 H, dd, *J*=12.5, 2.4 Hz), 7.41 - 7.59 (4 H, m), 7.14 - 7.33 (4 H, m), 6.42 (1 H, dd, *J*=5.3, 0.9 Hz), 4.52 (2 H, s), 3.97 (3 H, s), 2.63 - 2.72 (3 H, s), 2.04 - 2.10 (3 H, s)

## Example 116

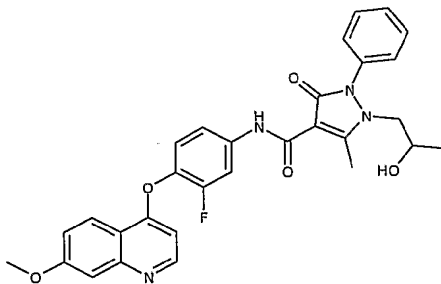


10

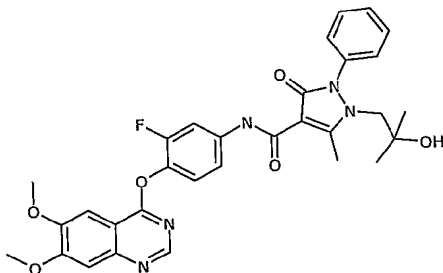
**1-(2,3-dihydroxy-2-methylpropyl)-N-(3-fluoro-4-((7-(methoxy)-4-quinolinyl)oxy)phenyl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) m/z: 573 (MH<sup>+</sup>). Calc'd exact mass for C<sub>31</sub>H<sub>29</sub>FN<sub>4</sub>O<sub>6</sub>: 572. <sup>1</sup>H NMR (400 MHz, *CHLOROFORM-d*) 10.85 (1 H, s), 8.56 (1 H, d, *J*=5.3 Hz), 8.27 (1 H, d, *J*=9.2 Hz), 7.90 (1 H, dd, *J*=12.5, 1.6 Hz), 7.54 (2 H, t, *J*=7.7 Hz), 7.38 - 7.47 (2 H, m), 7.14 - 7.32 (5 H, m), 6.42 (1 H, d, *J*=5.3 Hz), 4.09 - 4.19 (1 H, m), 3.96 (3 H, s), 3.86 (1 H, d, *J*=15.7 Hz), 3.34 (2 H, s), 2.89 (3 H, s), 1.07 (3 H, s).

15

## Example 117

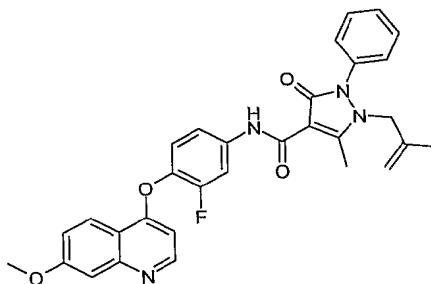


**N-(3-fluoro-4-((7-(methyloxy)-4-quinolinyl)oxy)phenyl)-1-(2-hydroxypropyl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) m/z: 543 (MH<sup>+</sup>). Calc'd exact mass for C<sub>28</sub>H<sub>25</sub>FN<sub>4</sub>O<sub>5</sub>: 542.

**Example 118**

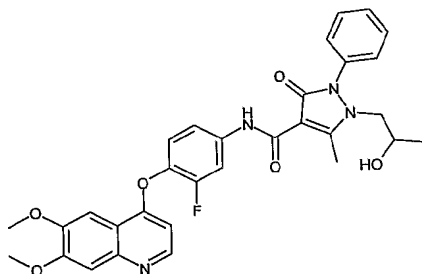
5

**N-(4-((6,7-bis(methyloxy)-4-quinazolinyl)oxy)-3-fluorophenyl)-1-(2-hydroxy-2-methylpropyl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) m/z: 588 (MH<sup>+</sup>). Calc'd exact mass for C<sub>31</sub>H<sub>30</sub>FN<sub>5</sub>O<sub>6</sub>: 587.

**Example 119**

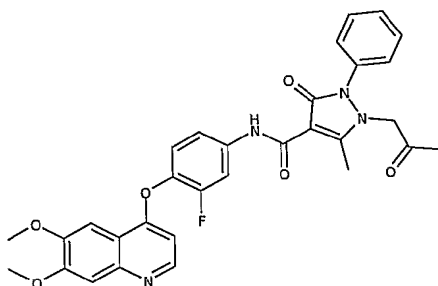
10

**N-(3-fluoro-4-((7-(methyloxy)-4-quinolinyl)oxy)phenyl)-5-methyl-1-(2-methyl-2-propen-1-yl)-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) m/z: 539 (MH<sup>+</sup>). Calc'd exact mass for C<sub>31</sub>H<sub>27</sub>FN<sub>4</sub>O<sub>4</sub>: 538.

**Example 120**

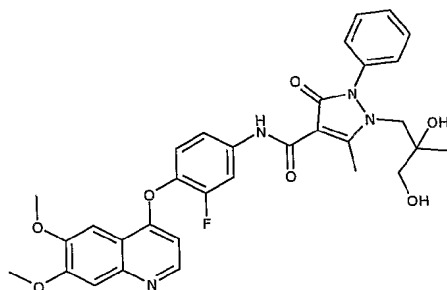
15

**N-(4-((6,7-bis(methyloxy)-4-quinolinyl)oxy)-3-fluorophenyl)-1-((2S)-2-hydroxypropyl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) m/z: 573 (MH<sup>+</sup>). Calc'd exact mass for C<sub>31</sub>H<sub>29</sub>FN<sub>4</sub>O<sub>6</sub>: 572.

**Example 121**

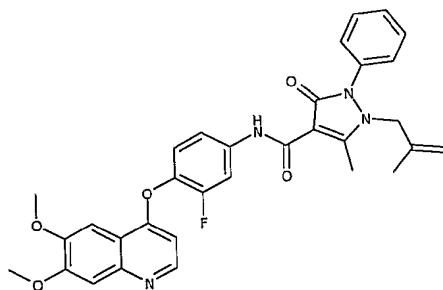
**N-(4-((6,7-bis(methoxy)-4-quinolinyl)oxy)-3-fluorophenyl)-5-methyl-3-oxo-1-(2-oxopropyl)-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) m/z:

5 571 (MH<sup>+</sup>). Calc'd exact mass for C<sub>31</sub>H<sub>27</sub>FN<sub>4</sub>O<sub>6</sub>: 570.

**Example 122**

**N-(4-((6,7-bis(methoxy)-4-quinolinyl)oxy)-3-fluorophenyl)-1-(2,3-dihydroxy-2-methylpropyl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS

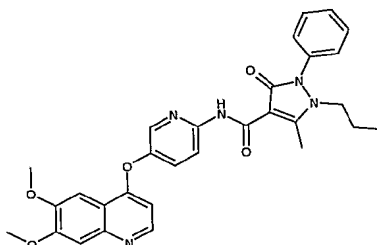
10 (ESI pos. ion) m/z: 603 (MH<sup>+</sup>). Calc'd exact mass for C<sub>32</sub>H<sub>31</sub>FN<sub>4</sub>O<sub>7</sub>: 602.

**Example 123**

**N-(4-((6,7-bis(methoxy)-4-quinolinyl)oxy)-3-fluorophenyl)-5-methyl-1-(2-methyl-2-propen-1-yl)-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)

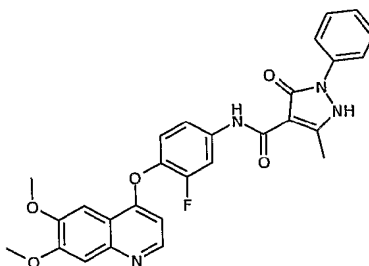
15 m/z: 569 (MH<sup>+</sup>). Calc'd exact mass for C<sub>32</sub>H<sub>29</sub>FN<sub>4</sub>O<sub>5</sub>: 568. <sup>1</sup>H NMR (400 MHz, MeOH) 10.16 (2 H, s), 7.73 (2 H, d, *J*=5.7 Hz), 7.24 (2 H, s), 7.21 (1 H, d, *J*=1.6 Hz), 6.94 (2 H, s), 6.80 - 6.90 (8 H, m), 6.58 - 6.71 (13 H, m), 5.86 (2 H, d, *J*=5.5 Hz), 4.13 - 4.19 (17 H, m), 3.65 - 3.72 (8 H, m), 3.30 (15 H, d, *J*=1.8 Hz), 2.57 - 2.60 (6 H, m), 2.27 (1 H, s), 1.99 - 2.14 (13 H, m), 0.84 (8 H, s)

## Example 124



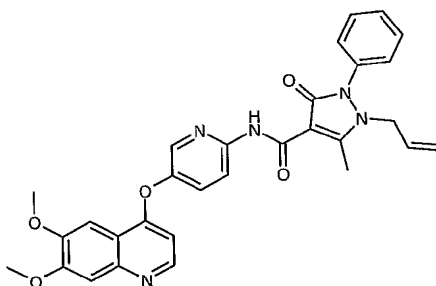
**N-(5-((6,7-bis(methoxy)-4-quinolinyloxy)-2-pyridinyl)-5-methyl-3-oxo-2-phenyl-1-propyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)  $m/z$ : 540 (MH<sup>+</sup>).  
 Calc'd exact mass for C<sub>30</sub>H<sub>29</sub>N<sub>5</sub>O<sub>5</sub>: 539.

## Example 125

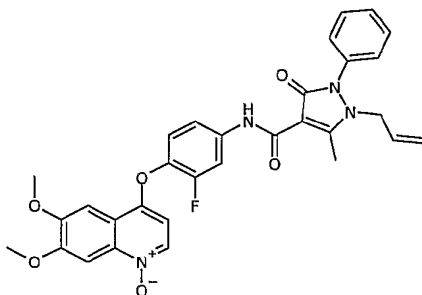


**N-(4-((6,7-bis(methoxy)-4-quinolinyloxy)-3-fluorophenyl)-5-methyl-3-oxo-2-phenyl-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)  $m/z$ : 515 (MH<sup>+</sup>). Calc'd exact mass for C<sub>28</sub>H<sub>23</sub>N<sub>4</sub>O<sub>5</sub>: 514.

## Example 126

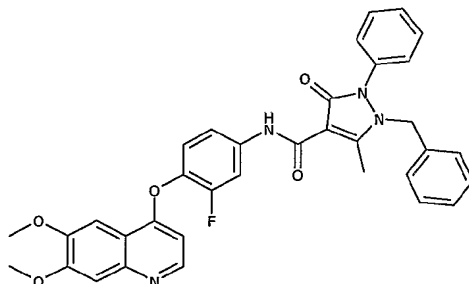


**N-(5-((6,7-bis(methoxy)-4-quinolinyloxy)-2-pyridinyl)-5-methyl-3-oxo-2-phenyl-1-(2-propen-1-yl)-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)  $m/z$ : 538 (MH<sup>+</sup>).  
 Calc'd exact mass for C<sub>30</sub>H<sub>27</sub>N<sub>5</sub>O<sub>5</sub>: 537.

**Example 127**

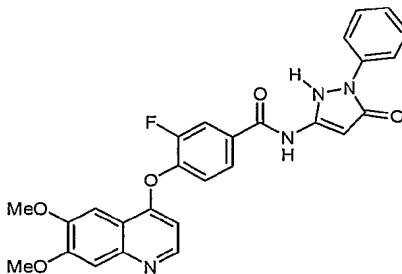
**N-(4-((6,7-bis(methoxy)-1-oxido-4-quinolinyl)oxy)-3-fluorophenyl)-5-methyl-3-oxo-2-phenyl-1-(2-propen-1-yl)-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion)

m/z: 571 (MH<sup>+</sup>). Calc'd exact mass for C<sub>31</sub>H<sub>27</sub>FN<sub>4</sub>O<sub>6</sub>: 570. <sup>1</sup>H NMR (400 MHz, MeOH) 8.36 (1 H, d, *J*=7.0 Hz), 7.90 - 7.97 (2 H, m), 7.68 (1 H, s), 7.53 - 7.62 (3 H, m), 7.42 (2 H, d, *J*=7.2 Hz), 7.30 - 7.37 (2 H, m), 6.59 (1 H, d, *J*=6.8 Hz), 5.69 (1 H, dd, *J*=11.2, 5.9 Hz), 5.21 (1 H, d, *J*=10.4 Hz), 4.93 (1 H, d, *J*=17.2 Hz), 4.48 (2 H, d, *J*=4.9 Hz), 4.01 - 4.11 (6 H, m), 2.77 (3 H, s).

**Example 128**

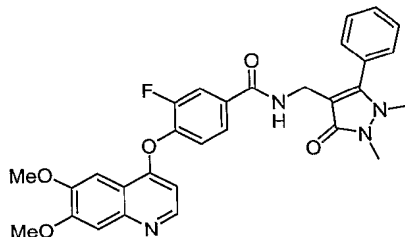
**N-(4-((6,7-bis(methoxy)-4-quinolinyl)oxy)-3-fluorophenyl)-5-methyl-3-oxo-2-phenyl-1-(phenylmethyl)-2,3-dihydro-1H-pyrazole-4-carboxamide:** MS (ESI pos. ion) m/z: 605

(MH<sup>+</sup>). Calc'd exact mass for C<sub>35</sub>H<sub>29</sub>FN<sub>4</sub>O<sub>5</sub>: 604.

**Example 129**

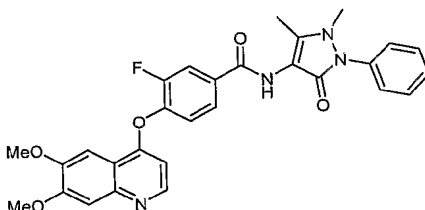
**4-(6,7-Dimethoxyquinolin-4-yloxy)-3-fluoro-N-(5-oxo-1-phenyl-2,5-dihydro-1H-pyrazol-3-yl)benzamide:** Calc'd for C<sub>27</sub>H<sub>21</sub>FN<sub>4</sub>O<sub>5</sub>: 500; MS (ESI pos. ion) m/z: 501 (MH<sup>+</sup>)

<sup>1</sup>HNMR (DMSO-d<sub>6</sub>, 400 MHz): 8.54 (1H, d, J 5.2), 7.88 (1H, dd, J 11.6, 1.6), 7.71 (1H, d, J 8.4), 7.59-7.54 (3H, m), 7.48 – 7.37 (4H, m), 7.13 (1H, t, J 7.3), 6.56 (1H, d, J 5.1), 3.96 (6H, s).

**Example 130**

**4-(6,7-Dimethoxyquinolin-4-yloxy)-N-((1,2-dimethyl-5-oxo-3-phenyl-2,5-dihydro-1H-pyrazol-4-yl)methyl)-3-fluorobenzamide:** Calc'd for C<sub>30</sub>H<sub>27</sub>FN<sub>4</sub>O<sub>5</sub>: 542; MS (ESI pos. ion) m/z: 543 (MH<sup>+</sup>)

<sup>1</sup>HNMR (CDCl<sub>3</sub>, 400 MHz): 8.51 (2H, m), 7.81 (1H, dd, J 1.8, 10.8), 7.71 (1H, d, J 8.3), 7.57-7.48 (5H, m), 7.44 (1H, s), 7.28 (1H, t, J 8.0), 6.42 (1H, d, J 5.3), 4.46 (d, 2H, d, J 4.9), 4.06 (6H, s), 3.48 (3H, s), 3.14 (3H, s).

**Example 131**

**4-(6,7-Dimethoxyquinolin-4-yloxy)-N-(2,3-dimethyl-5-oxo-1-phenyl-2,5-dihydro-1H-pyrazol-4-yl)-3-fluorobenzamide:** Calc'd for C<sub>29</sub>H<sub>25</sub>FN<sub>4</sub>O<sub>5</sub>: 528; MS (ESI pos. ion) m/z: 529 (MH<sup>+</sup>).

<sup>1</sup>HNMR (CDCl<sub>3</sub>, 400 MHz): 9.20 (1H, b), 8.56 (1H, d, J 5.3), 7.86 (1H, J), 7.76 (1H, d, J) 7.57 (1H, s), 7.50-7.44 (3H, m), 7.40 (2H, d, J 7.4), 7.34 (1H, t, J 7.2), 7.28 (1H, m), 6.56 (1H, d, J 4.9), 4.07 (3H, s), 4.06 (3H, s), 3.12 (3H, s), 2.30 (3H, s).

**Example 132**